

10542169.trn

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LOGINID : SSSPTA1626GMS

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

* * * * * * * * * * * * * Welcome to STN International * * * * * * * * * * * * *

NEWS 1 Web Page for STN Seminar Schedule - N. America
NEWS 2 AUG 06 CAS REGISTRY enhanced with new experimental property tags
NEWS 3 AUG 06 FSTA enhanced with new thesaurus edition
NEWS 4 AUG 13 CA/CAplus enhanced with additional kind codes for granted patents
NEWS 5 AUG 20 CA/CAplus enhanced with CAS indexing in pre-1907 records
NEWS 6 AUG 27 Full-text patent databases enhanced with predefined patent family display formats from INPADOCDB
NEWS 7 AUG 27 USPATOLD now available on STN
NEWS 8 AUG 28 CAS REGISTRY enhanced with additional experimental spectral property data
NEWS 9 SEP 07 STN AnaVist, Version 2.0, now available with Derwent World Patents Index
NEWS 10 SEP 13 FORIS renamed to SOFIS.
NEWS 11 SEP 13 INPADOCDB enhanced with monthly SDI frequency
NEWS 12 SEP 17 CA/CAplus enhanced with printed CA page images from 1967-1998
NEWS 13 SEP 17 CAplus coverage extended to include traditional medicine patents
NEWS 14 SEP 24 EMBASE, EMBAL, and LEMBASE reloaded with enhancements
NEWS 15 OCT 02 CA/CAplus enhanced with pre-1907 records from Chemisches Zentralblatt
NEWS 16 OCT 19 BEILSTEIN updated with new compounds
NEWS 17 NOV 15 Derwent Indian patent publication number format enhanced
NEWS 18 NOV 19 WPIX enhanced with XML display format
NEWS 19 NOV 30 ICSD reloaded with enhancements
NEWS 20 DEC 04 LINPADOCDB now available on STN
NEWS 21 DEC 14 BEILSTEIN pricing structure to change
NEWS 22 DEC 17 USPATOLD added to additional database clusters
NEWS 23 DEC 17 IMSDRUGCONF removed from database clusters and STN
NEWS 24 DEC 17 DGENE now includes more than 10 million sequences
NEWS 25 DEC 17 TOXCENTER enhanced with 2008 MeSH vocabulary in MEDLINE segment
NEWS 26 DEC 17 MEDLINE and LMEDLINE updated with 2008 MeSH vocabulary
NEWS 27 DEC 17 CA/CAplus enhanced with new custom IPC display formats
NEWS 28 DEC 17 STN Viewer enhanced with full-text patent content from USPATOLD
NEWS 29 JAN 02 STN pricing information for 2008 now available

NEWS EXPRESS 19 SEPTEMBER 2007: CURRENT WINDOWS VERSION IS V8.2, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 19 SEPTEMBER 2007.

NEWS HOURS STN Operating Hours Plus Help Desk Availability

Page 1

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NEWS LOGIN Welcome Banner and News Items
NEWS IPC8 For general information regarding STN implementation of IPC 8

Enter NEWS followed by the item number or name to see news on that specific topic.

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FILE 'HOME' ENTERED AT 07:29:54 ON 07 JAN 2008

```
=>
Uploading
THIS COMMAND NOT AVAILABLE IN THE CURRENT FILE
Do you want to switch to the Registry File?
Choice (Y/n) :
Switching to the Registry File...
Some commands only work in certain files. For
command can only be used to look at the index
index. Enter "HELP COMMANDS" at an arrow promp
commands which can be used in this file.
```

=> FILE REGISTRY

| COST IN U.S. DOLLARS | SINCE FILE
ENTRY | TOTAL
SESSION |
|----------------------|---------------------|------------------|
| FULL ESTIMATED COST | 0.21 | 0.21 |

FILE 'REGISTRY' ENTERED AT 07:30:13 ON 07 JAN 2008
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2008 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 6 JAN 2008 HIGHEST RN 960045-19-6
DICTIONARY FILE UPDATES: 6 JAN 2008 HIGHEST RN 960045-19-6

New CAS Information Use Policies - enter HELP-USAGETERMS for details

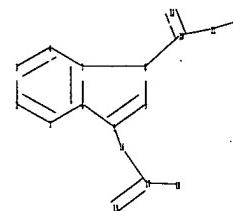
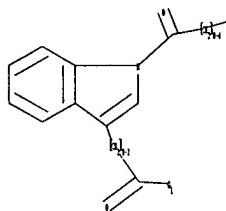
TSCA INFORMATION NOW CURRENT THROUGH June 29, 2007

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/support/stnqen/stndoc/properties.html>

=>
Uploading C:\Program Files\Stnexp\Queries\10542169.str



chain nodes :

10 11 12 13 16 17 18 19

ring nodes :

1 2 3 4 5 6 7 8 9

chain bonds :

5-16 8-10 10-11 10-12 11-13 16-17 17-18 17-19

ring bonds :

1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9

exact/norm bonds :

7-8 8-9 8-10 10-12 17-18 17-19

exact bonds :

5-6 5-9 5-16 10-11 11-13 16-17

normalized bonds :

1-2 1-6 2-3 3-4 4-7 6-7

isolated ring systems :

containing 1 :

G1:CH,O,N

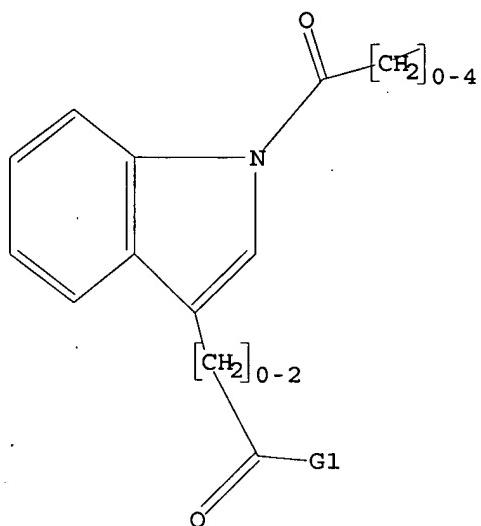
Match level :

| | | | | | | | | | |
|----------|----------|----------|----------|----------|----------|----------|--------|--------|----------|
| 1:Atom | 2:Atom | 3:Atom | 4:Atom | 5:Atom | 6:Atom | 7:Atom | 8:Atom | 9:Atom | 10:CLASS |
| 11:CLASS | 12:CLASS | 13:CLASS | 16:CLASS | 17:CLASS | 18:CLASS | 19:CLASS | | | |

L1 STRUCTURE UPLOADED

10542169.trn

=> d 11
L1 HAS NO ANSWERS
L1 STR



G1 CH₂O,N

Structure attributes must be viewed using STN Express query preparation.

=> s 11
SAMPLE SEARCH INITIATED 07:30:31 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 721 TO ITERATE

100.0% PROCESSED 721 ITERATIONS 34 ANSWERS
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 12810 TO 16030
PROJECTED ANSWERS: 331 TO 1029

L2 34 SEA SSS SAM L1

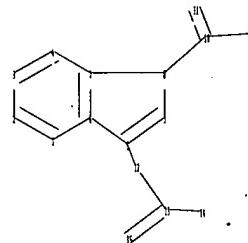
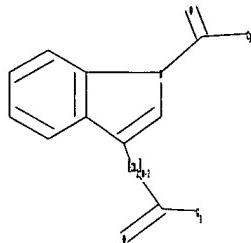
=> s 11 sss full
FULL SEARCH INITIATED 07:30:39 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 13810 TO ITERATE

100.0% PROCESSED 13810 ITERATIONS
SEARCH TIME: 00.00.01

L3 502 SEA SSS FUL L1

=>
Uploading C:\Program Files\Stnexp\Queries\10542169a.str

502 ANSWERS



chain nodes :

10 11 12 13 14 15 20

ring nodes :

1 2 3 4 5 6 7 8 9

chain bonds :

5-12 8-10 10-11 10-20 12-13 13-14 13-15

ring bonds :

1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9

exact/norm bonds :

7-8 8-9 8-10 10-11 10-20 13-14 13-15

exact bonds :

5-6 5-9 5-12 12-13

normalized bonds :

1-2 1-6 2-3 3-4 4-7 6-7

isolated ring systems :

containing 1 :

G1:CH,O,N

Match level :

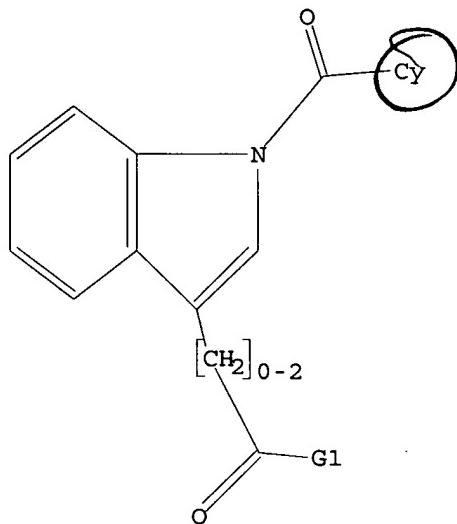
 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS
 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 20:CLASS

L4 STRUCTURE UPLOADED

=> d 14

L4 HAS NO ANSWERS

L4 STR



G1 CH₂O,N

Structure attributes must be viewed using STN Express query preparation.

```
=> s 14
SAMPLE SEARCH INITIATED 07:34:51 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 961 TO ITERATE
```

```
100.0% PROCESSED 961 ITERATIONS
INCOMPLETE SEARCH (SYSTEM LIMIT EXCEEDED)
SEARCH TIME: 00.00.01
```

```
FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 17361 TO 21079
PROJECTED ANSWERS: 2973 TO 4627
```

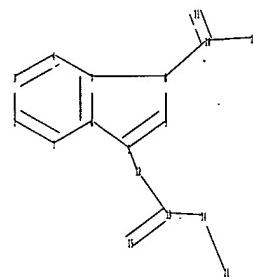
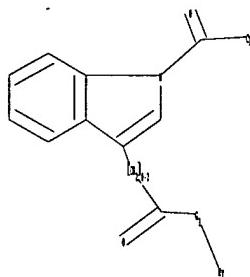
L5 50 SEA SSS SAM L4

```
=> s 14 sss full
FULL SEARCH INITIATED 07:35:04 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 18894 TO ITERATE
```

```
100.0% PROCESSED 18894 ITERATIONS
SEARCH TIME: 00.00.01
```

L6 3642 SEA SSS FUL L4

```
=>
Uploading C:\Program Files\Stnexp\Queries\10542169b.str
```



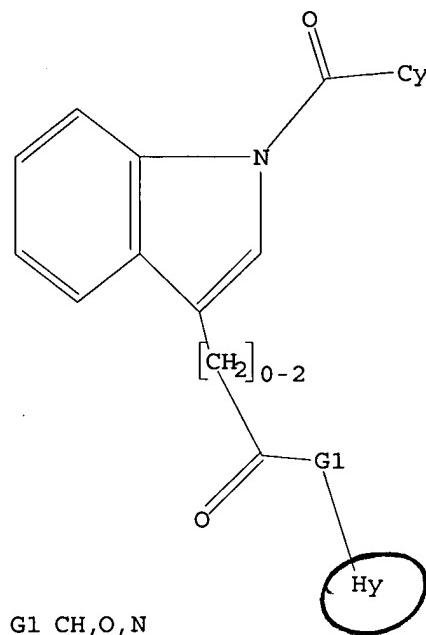
chain nodes :
 10 11 12 13 14 15 20 21
 ring nodes :
 1 2 3 4 5 6 7 8 9
 chain bonds :
 5-12 8-10 10-11 10-20 12-13 13-14 13-15 14-21
 ring bonds :
 1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9
 exact/norm bonds :
 7-8 8-9 8-10 10-11 10-20 13-14 13-15 14-21
 exact bonds :
 5-6 5-9 5-12 12-13
 normalized bonds :
 1-2 1-6 2-3 3-4 4-7 6-7
 isolated ring systems :
 containing 1 :

G1:CH,O,N

Match level :
 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS
 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 20:CLASS 21:Atom

L7 STRUCTURE UPLOADED

=> d 17
 L7 HAS NO ANSWERS
 L7 STR



Structure attributes must be viewed using STN Express query preparation.

=> s 17
SAMPLE SEARCH INITIATED 07:36:56 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 961 TO ITERATE

100.0% PROCESSED 961 ITERATIONS 11 ANSWERS
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 17361 TO 21079
PROJECTED ANSWERS: 22 TO 418

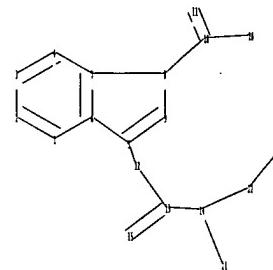
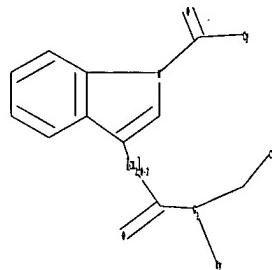
L8 11 SEA SSS SAM L7

=> s 17 sss full
FULL SEARCH INITIATED 07:37:03 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 18894 TO ITERATE

100.0% PROCESSED 18894 ITERATIONS 215 ANSWERS
SEARCH TIME: 00.00.01

L9 215 SEA SSS FUL L7

=>
Uploading C:\Program Files\Stnexp\Queries\10542169c.str



chain nodes :

10 11 12 13 14 15 20 21 22 23

ring nodes :

1 2 3 4 5 6 7 8 9

chain bonds :

5-12 8-10 10-11 10-20 12-13 13-14 13-15 14-21 14-22 22-23

ring bonds :

1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9

exact/norm bonds :

7-8 8-9 8-10 10-11 10-20 13-14 13-15 14-21 14-22

exact bonds :

5-6 5-9 5-12 12-13 22-23

normalized bonds :

1-2 1-6 2-3 3-4 4-7 6-7

isolated ring systems :

containing 1 :

G1:CH,O,N

Match level :

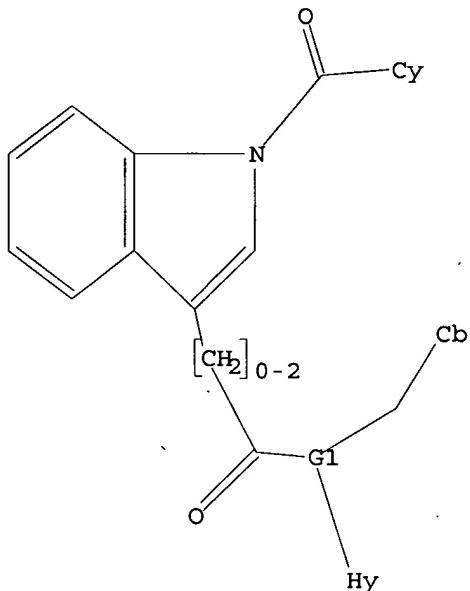
 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS
 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 20:CLASS 21:Atom 22:CLASS
 23:Atom

L10 STRUCTURE UPLOADED

=> d 110

L10 HAS NO ANSWERS

L10 STR

G1 CH₂O,N

Structure attributes must be viewed using STN Express query preparation.

```
=> s 110
SAMPLE SEARCH INITIATED 07:39:54 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 961 TO ITERATE
```

```
100.0% PROCESSED 961 ITERATIONS 3 ANSWERS
SEARCH TIME: 00.00.01
```

```
FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 17361 TO 21079
PROJECTED ANSWERS: 3 TO 163
```

L11 3 SEA SSS SAM L10

```
=> s 110 sss full
FULL SEARCH INITIATED 07:40:01 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 18863 TO ITERATE
```

```
100.0% PROCESSED 18863 ITERATIONS
SEARCH TIME: 00.00.01
```

14 ANSWERS

L12 14 SEA SSS FUL L10

| | | |
|----------------------|------------|---------|
| => FIL HCAPLUS | SINCE FILE | TOTAL |
| COST IN U.S. DOLLARS | ENTRY | SESSION |
| FULL ESTIMATED COST | 718.96 | 719.17 |

```
FILE 'HCAPLUS' ENTERED AT 07:40:06 ON 07 JAN 2008
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FILE COVERS 1907 - 7 Jan 2008 VOL 148 ISS 2
FILE LAST UPDATED: 6 Jan 2008 (20080106/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d his

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FILE 'REGISTRY' ENTERED AT 07:30:13 ON 07 JAN 2008

| | | | |
|-----|------|-----------|-------------|
| L1 | | STRUCTURE | UPLOADED |
| L2 | 34 | S L1 | |
| L3 | 502 | S L1 SSS | FULL |
| L4 | | STRUCTURE | UPLOADED |
| L5 | 50 | S L4 | |
| L6 | 3642 | S L4 SSS | <u>FULL</u> |
| L7 | | STRUCTURE | UPLOADED |
| L8 | 11 | S L7 | |
| L9 | 215 | S L7 SSS | FULL |
| L10 | | STRUCTURE | UPLOADED |
| L11 | 3 | S L10 | |
| L12 | 14 | S L10 SSS | FULL |

FILE 'HCAPLUS' ENTERED AT 07:40:06 ON 07 JAN 2008

=> s 16
L13 16343 L6

=> S 19
L14 113 L9

=> s 112
L15 1 L12

=> s 114 and ophthalmic
11855 OPHTHALMIC
52 OPHTHALMICS
11876 OPHTHALMIC
(OPHTHALMIC OR OPHTHALMICS)

=> s 114 and ocular hypertension
19018 OCULAR
22 OCULARS

10542169.trn

19036 OCULAR
(OCULAR OR OCULARS)
91866 HYPERTENSION
106 HYPERTENSIONS
91887 HYPERTENSION
(HYPERTENSION OR HYPERTENSIONS)
933 OCULAR HYPERTENSION
(OCULAR (W) HYPERTENSION)

L17 1 L14 AND OCULAR HYPERTENSION

=> s l14 and hypertension
91866 HYPERTENSION
106 HYPERTENSIONS
91887 HYPERTENSION
(HYPERTENSION OR HYPERTENSIONS)

L18 1 L14 AND HYPERTENSION

=> s l14 and py<=2003
23975204 PY<=2003
L19 87 L14 AND PY<=2003

=> s l19 and p/dt
6030773 P/DT
L20 44 L19 AND P/DT

=> s l20 and us/pc
1756160 US/PC
L21 23 L20 AND US/PC

=> s l21 and treating
455033 TREATING
1 TREATINGS
455034 TREATING
(TREATING OR TREATINGS)

L22 4 L21 AND TREATING

=> s l21 and treat
82039 TREAT
9796 TREATS
91334 TREAT
(TREAT OR TREATS)

L23 1 L21 AND TREAT

=> s l21 and treatment
2375898 TREATMENT
222073 TREATMENTS
2492988 TREATMENT
(TREATMENT OR TREATMENTS)

L24 10 L21 AND TREATMENT

=> d his

(FILE 'HOME' ENTERED AT 07:29:54 ON 07 JAN 2008)

FILE 'REGISTRY' ENTERED AT 07:30:13 ON 07 JAN 2008

L1 STRUCTURE uploaded
L2 34 S L1
L3 502 S L1 SSS FULL
L4 STRUCTURE uploaded
L5 50 S L4

10542169.trn

L6 3642 S L4 SSS FULL
L7 STRUCTURE uploaded
L8 11 S L7
L9 215 S L7 SSS FULL
L10 STRUCTURE uploaded
L11 3 S L10
L12 14 S L10 SSS FULL

FILE 'HCAPLUS' ENTERED AT 07:40:06 ON 07 JAN 2008

L13 16343 S L6
L14 113 S L9
L15 1 S L12
L16 2 S L14 AND OPHTHALMIC
L17 1 S L14 AND OCULAR HYPERTENSION
L18 1 S L14 AND HYPERTENSION
L19 87 S L14 AND PY<=2003
L20 44 S L19 AND P/DT
L21 23 S L20 AND US/PC
L22 4 S L21 AND TREATING
L23 1 S L21 AND TREAT
L24 10 S L21 AND TREATMENT

=> d 115 ibib abs hitstr tot

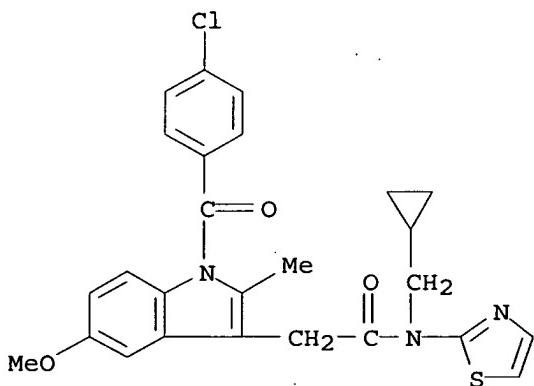
L15 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER: 2004:857325 HCAPLUS
DOCUMENT NUMBER: 141:350033
TITLE: Preparation of 5-methoxy-2-methylindole-3-acetamide derivs. as potassium channel blockers for treating ocular hypertension
INVENTOR(S): Fisher, Michael H.; Garcia, Maria L.; Kaczorowski, Gregory J.; Meinke, Peter T.; Parsons, William H.; Boyd, Edward Andrew; Price, Stephen; Stibbard, John
PATENT ASSIGNEE(S): Merck & Co., Inc., USA; Evotec Oai
SOURCE: PCT Int. Appl., 109 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|----------|
| WO 2004087051 | A2 | 20041014 | WO 2004-US9028 | 20040324 |
| WO 2004087051 | A3 | 20050721 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
TD, TG | | | | |
| AU 2004226479 | A1 | 20041014 | AU 2004-226479 | 20040324 |
| CA 2519899 | A1 | 20041014 | CA 2004-2519899 | 20040324 |
| EP 1610776 | A2 | 20060104 | EP 2004-758273 | 20040324 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, | | | | |

| | | | | |
|--|--|----------|------------------|------------|
| IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK | | | | |
| CN 1791402 | A | 20060621 | CN 2004-80013916 | 20040324 |
| JP 2006524239 | T | 20061026 | JP 2006-509260 | 20040324 |
| US 2006069256 | Al | 20060330 | US 2005-542169 | 20050713 |
| IN 2005DN04100 | A | 20070831 | IN 2005-DN4100 | 20050912 |
| PRIORITY APPLN. INFO.: | | | US 2003-458103P | P 20030327 |
| | | | WO 2004-US9028 | A 20040324 |
| OTHER SOURCE(S) : | CASREACT 141:350033; MARPAT 141:350033 | | | |
| GI | | | | |

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

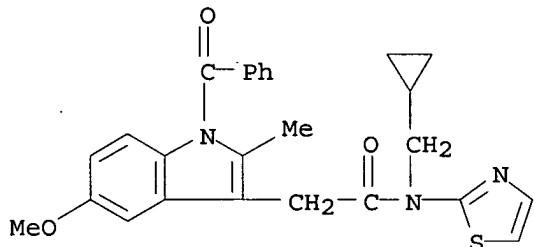
- AB The title compds. I [X = -(CHR₇)p-; Y = -CO(CH₂)_n- or -CH(OR₈)-; Q = N, CR₉, or O; R₁ = H, alkyl, CF₃, alkoxy, OH, etc.; R₂ = H, alkyl, alkylSR₈, -(CH₂)_nO(CH₂)_mOR₈, -(CH₂)alkoxy, etc.; R₃ = H, alkyl, -(CH₂)ncycloalkyl, -(CH₂)nheterocyclyl, or when Q = N, R₂, R₃ taken together with the the N form a 4-10 membered heterocyclic ring; R₄, R₅ = H, alkoxy, OH, alkyl, COOR₈, SO₃H, etc.; R₆ = H, alkyl, -(CH₂)(hetero)aryl, -NH(CH₂)(hetero)aryl, etc.; R₇ = H, alkyl, -(CH₂)nCOOR₈, or -(CH₂)nN(R₈)₂; R₈ = H, or alkyl; R₉ = H, or alkyl; m = 0-3; n = 0-3, p = 0-1] were prepared as potent potassium channel blockers in the treatment of glaucoma and other conditions which leads to elevated intraocular pressure in the eye of a patient. For example, reaction of 1-(4-chlorobenzoyl)-5-methoxy-2-methylindole-3-acetic acid with N-cyclohexyl-N-thiazol-2-yl amine (preparation given) yielded compound II. The compds. of this invention inhibited Maxi-K Channel activity with IC₅₀'s in the range of 1 nM to 20 μM.
- IT 773898-40-1P 773898-77-4P 773898-78-5P
 773898-79-6P 773898-80-9P 773898-81-0P
 773898-82-1P 773898-83-2P 773898-91-2P
 773898-92-3P 773898-93-4P 773898-94-5P
 773898-97-8P 773898-98-9P
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation of 5-methoxy-2-methylindole-3-acetamide derivs. as potassium channel blockers for treating ocular hypertension)
- RN 773898-40-1 HCPLUS
 CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



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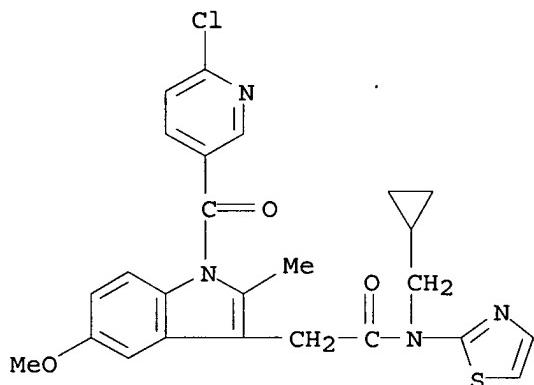
RN 773898-77-4 HCPLUS

CN 1H-Indole-3-acetamide, 1-benzoyl-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



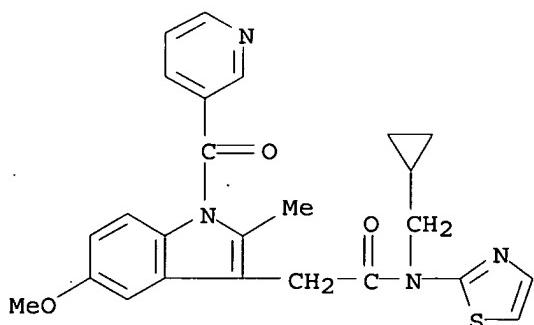
RN 773898-78-5 HCPLUS

CN 1H-Indole-3-acetamide, 1-[(6-chloro-3-pyridinyl)carbonyl]-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



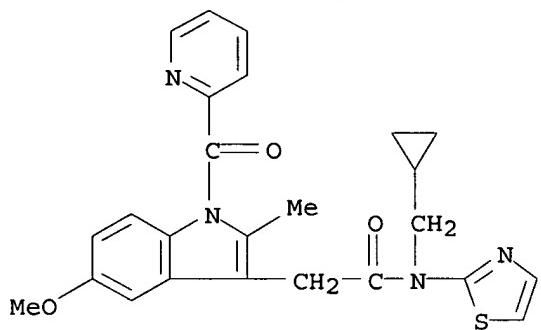
RN 773898-79-6 HCPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(3-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



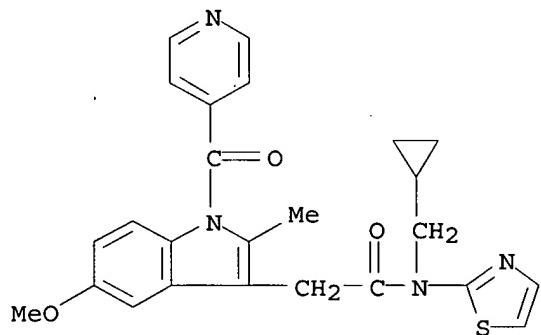
RN 773898-80-9 HCPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(2-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



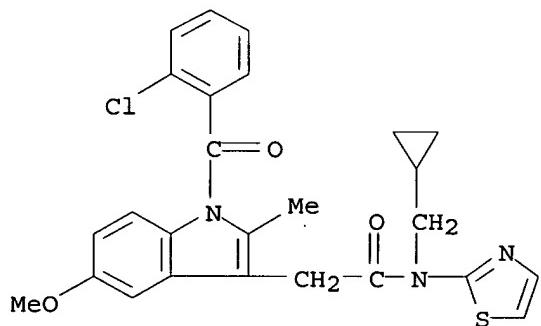
RN 773898-81-0 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(4-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



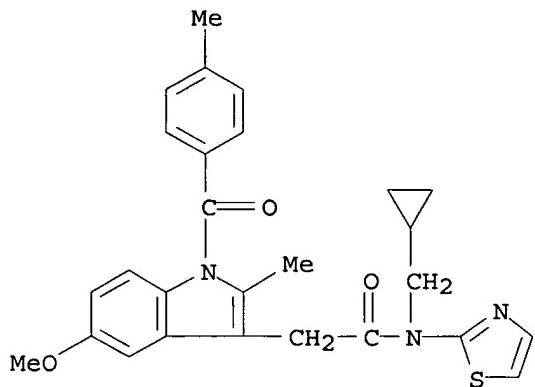
RN 773898-82-1 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(2-chlorobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)

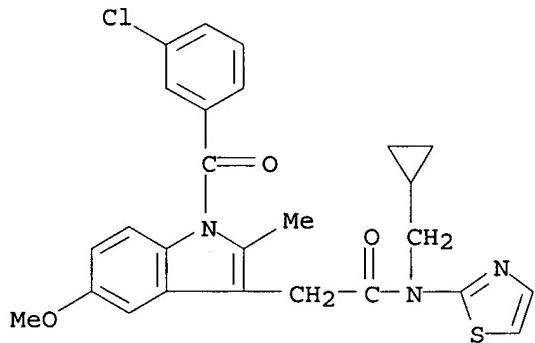


RN 773898-83-2 HCAPLUS

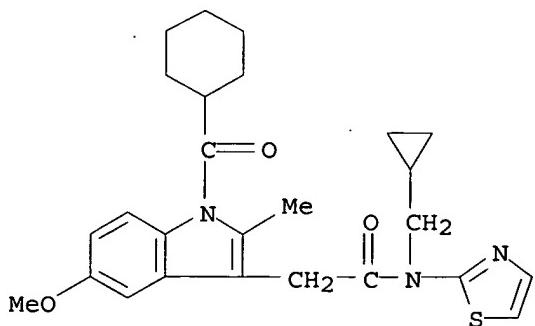
CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(4-methylbenzoyl)-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-91-2 HCPLUS
CN 1H-Indole-3-acetamide, 1-(3-chlorobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)

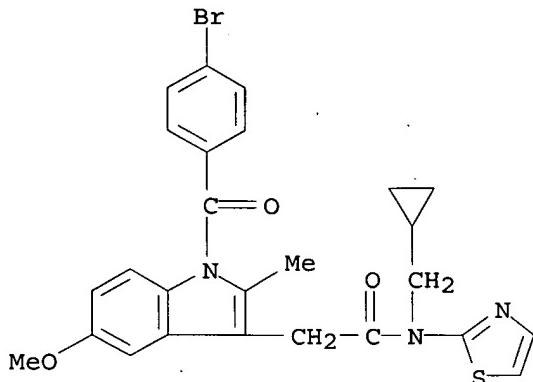


RN 773898-92-3 HCPLUS
CN 1H-Indole-3-acetamide, 1-(cyclohexylcarbonyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



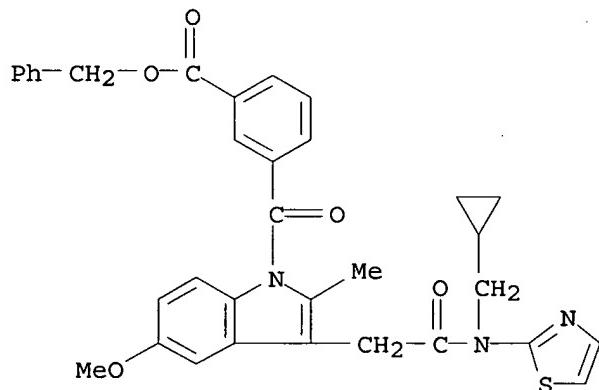
RN 773898-93-4 HCPLUS
CN 1H-Indole-3-acetamide, 1-(4-bromobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)

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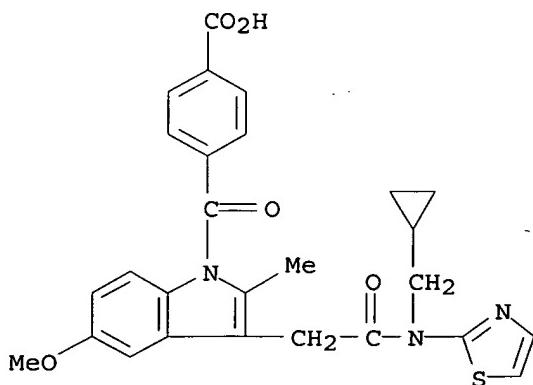
RN 773898-94-5 HCAPLUS

CN Benzoic acid, 3-[[3-[(cyclopropylmethyl)-2-thiazolylamino]-2-oxoethyl]-5-methoxy-2-methyl-1H-indol-1-yl]carbonyl-, phenylmethyl ester (CA INDEX NAME)



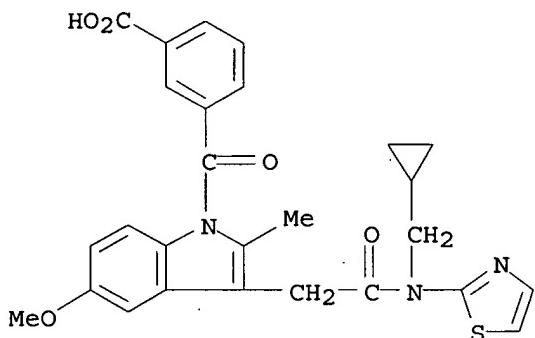
RN 773898-97-8 HCAPLUS

CN Benzoic acid, 4-[[3-[(cyclopropylmethyl)-2-thiazolylamino]-2-oxoethyl]-5-methoxy-2-methyl-1H-indol-1-yl]carbonyl- (CA INDEX NAME)



RN 773898-98-9 HCAPLUS

CN Benzoic acid, 3-[[3-[(cyclopropylmethyl)-2-thiazolylamino]-2-oxoethyl]-5-methoxy-2-methyl-1H-indol-1-yl]carbonyl] - (CA INDEX NAME)



=> d 116 ibib abs hitstr tot

L16 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2001:721438 HCAPLUS

DOCUMENT NUMBER: 135:288343

TITLE: Preparation and activity of nitrosated and
nitrosylated nonsteroidal antiinflammatory compounds

INVENTOR(S): Bandarage, Upul K.; Dong, Qing; Fang, Xinqin; Garvey,
David S.; Mercer, Gregory J.; Richardson, Stewart K.;
Schroeder, Joseph D.; Wang, Tiansheng

PATENT ASSIGNEE(S): Nitromed, Inc., USA

SOURCE: U.S., 59 pp., Cont.-in-part of U.S. Ser. No. 182,433,
abandoned.

CODEN: USXXAM

DOCUMENT TYPE: Patent

LANGUAGE: English

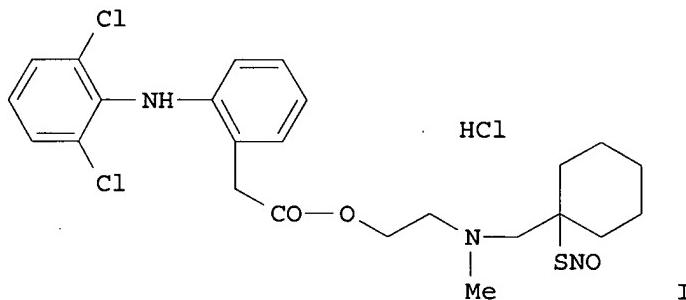
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|---------------------|----------|
| US 6297260 | B1 | 20011002 | US 1999-429019 | 19991029 |
| CA 2348741 | A1 | 20000511 | CA 1999-2348741 | 19991029 |
| WO 2000025776 | A1 | 20000511 | WO 1999-US25481 | 19991029 |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW | | | AM, AZ, BY, KG, KZ, | |
| MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE,
IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML,
MR, NE, SN, TD, TG | | | | |
| EP 1126838 | A1 | 20010829 | EP 1999-958708 | 19991029 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO | | | | |
| JP 2002528495 | T | 20020903 | JP 2000-579217 | 19991029 |
| AU 763000 | B2 | 20030710 | AU 2000-16012 | 19991029 |
| US 2002016322 | A1 | 20020207 | US 2001-938560 | 20010827 |
| US 6593347 | B2 | 20030715 | | |

| | | | | |
|------------------------|----|----------|-----------------|-------------|
| US 2003207919 | A1 | 20031106 | US 2003-431457 | 20030508 |
| AU 2004200091 | A1 | 20040205 | AU 2004-200091 | 20040109 |
| PRIORITY APPLN. INFO.: | | | US 1998-182433 | B2 19981030 |
| | | | AU 2000-16012 | A 19991029 |
| | | | US 1999-429019 | A3 19991029 |
| | | | WO 1999-US25481 | W 19991029 |
| | | | US 2001-938560 | A3 20010827 |

OTHER SOURCE(S) : MARPAT 135:288343
GI



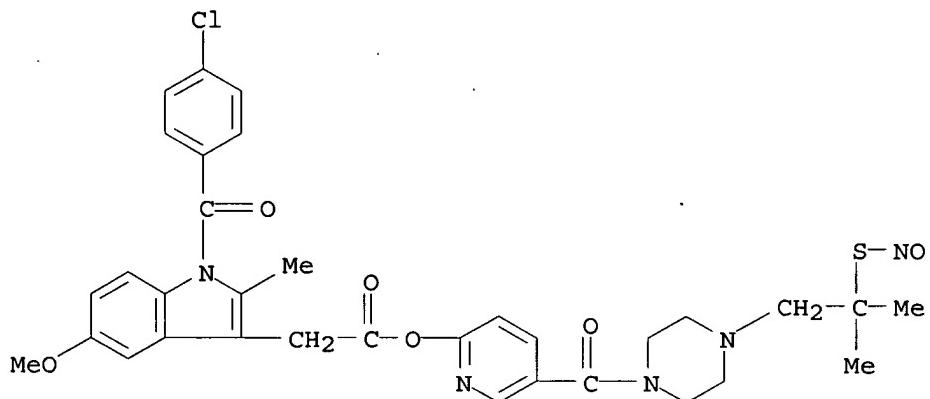
AB The present invention describes novel nitrosated and/or nitrosylated nonsteroidal antiinflammatory compds., and novel compns. comprising at least one nitrosated and/or nitrosylated nonsteroidal antiinflammatory compound, and, optionally, at least one compound that donates, transfers or releases nitric oxide, elevates endogenous levels of endothelium-derived relaxing factor, stimulates endogenous synthesis of nitric oxide or is a substrate for nitric oxide synthase. The present invention also provides methods for treating, preventing and/or reducing inflammation, pain, and fever; decreasing or reversing the gastrointestinal, renal and other toxicities resulting from the use of nonsteroidal antiinflammatory drugs; treating and/or preventing gastrointestinal disorders; treating inflammatory disease states and disorders; and treating and/or preventing ophthalmic diseases or disorders. Thus, I was prepared in 8 steps from cyclohexanecarboxaldehyde and shows a relative activity of 1, 1.2 and 0.02 in analgesic, antiinflammatory and gastric lesion tests.

IT 364590-30-7P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation and activity of nitrosated and nitrosylated nonsteroidal antiinflammatory compds.)

RN 364590-30-7 HCPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-, 5-[[4-[2-methyl-2-(nitrosothio)propyl]-1-piperazinyl]carbonyl]-2-pyridinyl ester (CA INDEX NAME)



REFERENCE COUNT: 63 THERE ARE 63 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L16 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 1995:858705 HCAPLUS
 DOCUMENT NUMBER: 123:266118
 TITLE: Codrugs as a method of controlled drug delivery
 INVENTOR(S): Ashton, Paul; Crooks, Peter Anthony; Riggs, Robert
 Mack; Cynkowski, Tadeusz; Cynkowska, Grazyna
 PATENT ASSIGNEE(S): University of Kentucky Research Foundation, USA
 SOURCE: PCT Int. Appl., 57 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO.. | KIND | DATE | APPLICATION NO. | DATE |
|---|--------------------|----------|--|---------------------------|
| WO 9520567 | A1 | 19950803 | WO 1994-US1659 | 19940217 |
| W: AU, CA, JP
RW: AT, BE, CH, CA 2182228 | DE, DK, ES, FR, A1 | 19950803 | GB, GR, IE, IT, LU, MC, NL, PT, SE
CA 1994-2182228 | 19940217 |
| AU 9462545 | A | 19950815 | AU 1994-62545 | 19940217 |
| AU 705226 | B2 | 19990520 | | |
| EP 740650 | A1 | 19961106 | EP 1994-909643 | 19940217 |
| EP 740650 | B1 | 20040526 | | |
| R: AT, BE, CH, JP 09509151 | DE, DK, ES, FR, T | 19970916 | GB, GR, IE, IT, LI, LU, MC, NL, PT, SE
JP 1994-520023 | 19940217 |
| AT 267798 | T | 20040615 | AT 1994-909643 | 19940217 |
| PT 740650 | T | 20041029 | PT 1994-909643 | 19940217 |
| ES 2222455 | T3 | 20050201 | ES 1994-909643 | 19940217 |
| US 6051576 | A | 20000418 | US 1997-791071
US 1994-187462 | 19970129
A 19940128 |
| PRIORITY APPLN. INFO.: | | | WO 1994-US1659
US 1995-388855 | W 19940217
B1 19950215 |

AB A codrug composition of at least two drug compds. covalently linked to one another via a labile bond to form a single codrug composition, and methods of use of the codrug for the treatment of various medical conditions are disclosed. The codrug may be administered by itself or as a bioerodible or nonbioerodible dosage form, such as injection, liposome, suspension, microsphere, nanoparticle, ointment, transdermal patch, etc.

IT 169046-88-2P

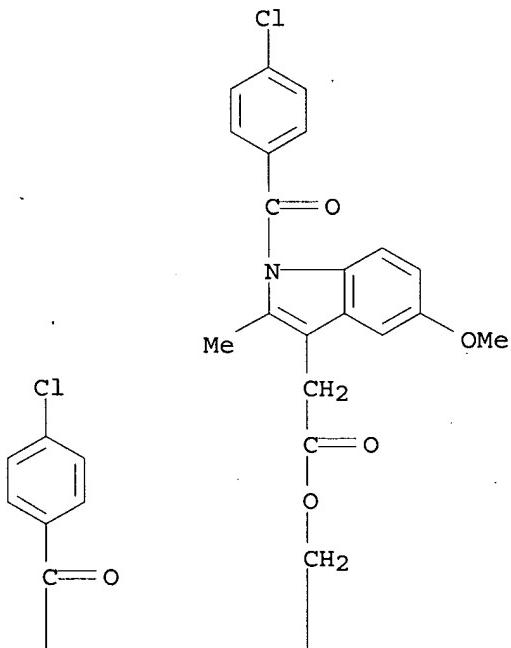
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RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(codrug compns. for controlled drug delivery)

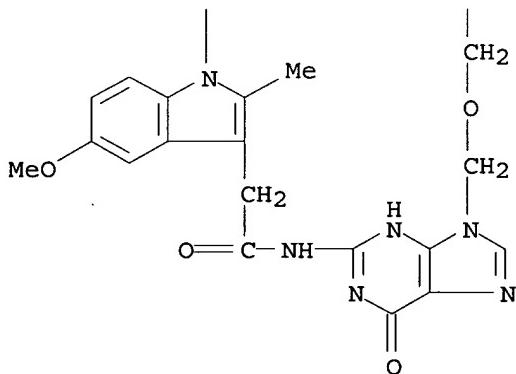
RN 169046-88-2 HCAPLUS

CN 1H-Indole-3-acetic acid, 1- (4-chlorobenzoyl)-5-methoxy-2-methyl-,
2- [[2- [[1- (4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-
yl]acetyl]amino]-1,6-dihydro-6-oxo-9H-purin-9-yl]methoxy]ethyl ester (9CI)
(CA INDEX NAME)

PAGE 1-A



PAGE 2-A



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L17 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2008 ACS ON STN
 ACCESSION NUMBER: 2004:857325 HCAPLUS
 DOCUMENT NUMBER: 141:350033
 TITLE: Preparation of 5-methoxy-2-methylindole-3-acetamide
 derivs. as potassium channel blockers for treating
 ocular hypertension
 INVENTOR(S): Fisher, Michael H.; Garcia, Maria L.; Kaczorowski,
 Gregory J.; Meinke, Peter T.; Parsons, William H.;
 Boyd, Edward Andrew; Price, Stephen; Stibbard, John
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA; Evotec Oai
 SOURCE: PCT Int. Appl., 109 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|------------------|------------|
| WO 2004087051 | A2 | 20041014 | WO 2004-US9028 | 20040324 |
| WO 2004087051 | A3 | 20050721 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
SK, TR, BF, BJ, CF, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
TD, TG | | | | |
| AU 2004226479 | A1 | 20041014 | AU 2004-226479 | 20040324 |
| CA 2519899 | A1 | 20041014 | CA 2004-2519899 | 20040324 |
| EP 1610776 | A2 | 20060104 | EP 2004-758273 | 20040324 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK | | | | |
| CN 1791402 | A | 20060621 | CN 2004-80013916 | 20040324 |
| JP 2006524239 | T | 20061026 | JP 2006-509260 | 20040324 |
| US 2006069256 | A1 | 20060330 | US 2005-542169 | 20050713 |
| IN 2005DN04100 | A | 20070831 | IN 2005-DN4100 | 20050912 |
| PRIORITY APPLN. INFO.: | | | US 2003-458103P | P 20030327 |
| | | | WO 2004-US9028 | A 20040324 |

OTHER SOURCE(S): CASREACT 141:350033; MARPAT 141:350033
 GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The title compds. I [X = -(CHR7)p-; Y = -CO(CH2)n- or -CH(OR8)-; Q = N, CR9, or O; R1 = H, alkyl, CF3, alkoxy, OH, etc.; R2 = H, alkyl, alkylSR8, -(CH2)nO(CH2)mOR8, -(CH2)alkoxy, etc.; R3 = H, alkyl, -(CH2)ncycloalkyl, -(CH2)nheterocyclyl, or when Q = N, R2, R3 taken together with the the N form a 4-10 membered heterocyclic ring; R4, R5 = H, alkoxy, OH, alkyl, COOR8, SO3H, etc.; R6 = H, alkyl, -(CH2)(hetero)aryl, -NH(CH2)(hetero)aryl, etc.; R7 = H, alkyl, -(CH2)nCOOR8, or -(CH2)nN(R8)2; R8 = H, or alkyl; R9 = H, or alkyl; m = 0-3; n = 0-3, p = 0-1] were prepared as potent potassium channel blockers in the treatment of glaucoma and

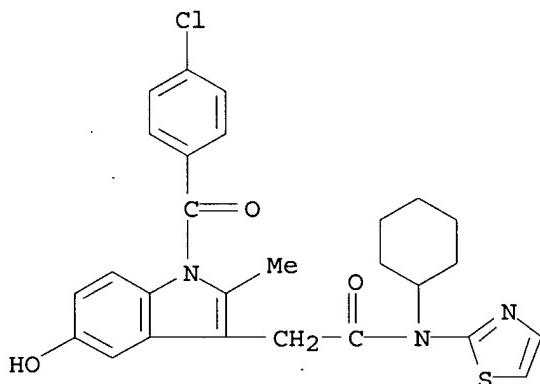
other conditions which leads to elevated intraocular pressure in the eye of a patient. For example, reaction of 1-(4-chlorobenzoyl)-5-methoxy-2-methylindole-3-acetic acid with N-cyclohexyl-N-thiazol-2-yl amine (preparation given) yielded compound II. The compds. of this invention inhibited Maxi-K Channel activity with IC₅₀'s in the range of 1 nM to 20 μM.

IT 773898-89-8P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
(preparation of 5-methoxy-2-methylindole-3-acetamide derivs. as potassium channel blockers for treating ocular hypertension)

RN 773898-89-8 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-5-hydroxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



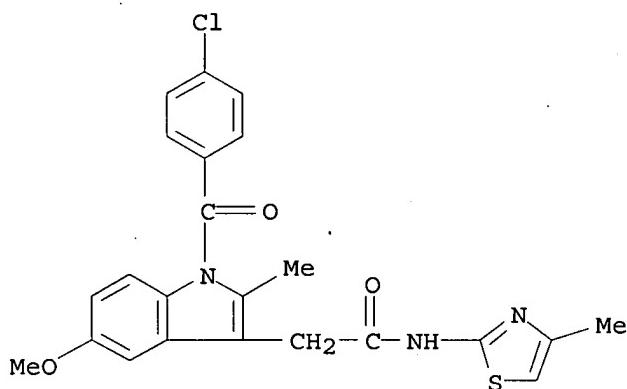
IT 282728-83-0P 732255-43-5P 773898-05-8P
773898-10-5P 773898-14-9P 773898-15-0P
773898-16-1P 773898-17-2P 773898-18-3P
773898-19-4P 773898-32-1P 773898-33-2P
773898-35-4P 773898-36-5P 773898-37-6P
773898-38-7P 773898-39-8P 773898-40-1P
773898-41-2P 773898-42-3P 773898-43-4P
773898-44-5P 773898-45-6P 773898-46-7P
773898-47-8P 773898-48-9P 773898-49-0P
773898-50-3P 773898-51-4P 773898-52-5P
773898-53-6P 773898-54-7P 773898-55-8P
773898-56-9P 773898-57-0P 773898-58-1P
773898-59-2P 773898-60-5P 773898-61-6P
773898-62-7P 773898-63-8P 773898-65-0P
773898-66-1P 773898-67-2P 773898-69-4P
773898-70-7P 773898-71-8P 773898-72-9P
773898-73-0P 773898-75-2P 773898-77-4P
773898-78-5P 773898-79-6P 773898-80-9P
773898-81-0P 773898-82-1P 773898-83-2P
773898-84-3P 773898-85-4P 773898-86-5P
773898-87-6P 773898-88-7P 773898-90-1P
773898-91-2P 773898-92-3P 773898-93-4P
773898-94-5P 773898-97-8P 773898-98-9P
773898-99-0P 773899-00-6P 773899-01-7P
773899-02-8P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of 5-methoxy-2-methylindole-3-acetamide derivs. as potassium channel blockers for treating ocular hypertension)

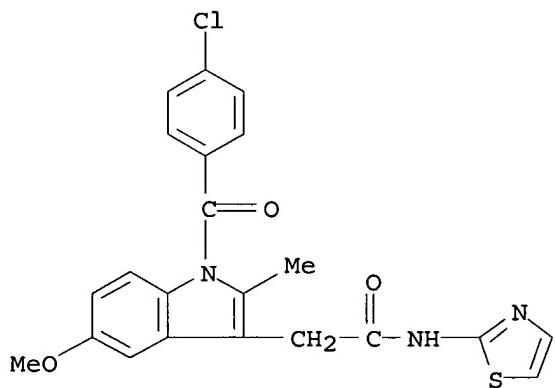
RN 282728-83-0 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(4-methyl-2-thiazolyl)- (CA INDEX NAME)



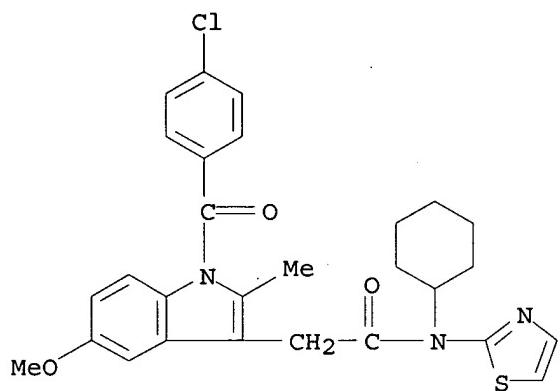
RN 732255-43-5 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



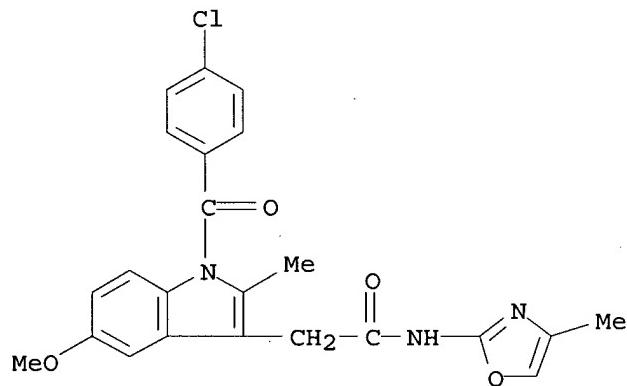
RN 773898-05-8 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



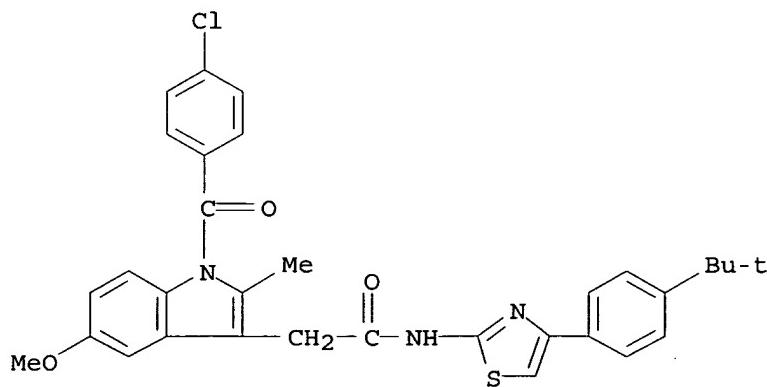
RN 773898-10-5 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(4-methyl-2-oxazolyl)- (CA INDEX NAME)



RN 773898-14-9 HCPLUS

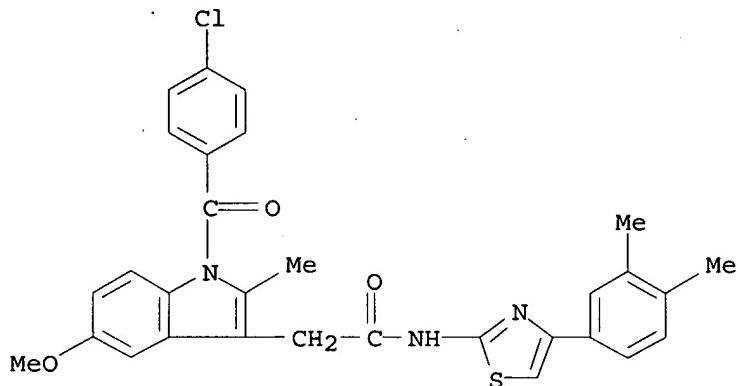
CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-[4-[4-(1,1-dimethylethyl)phenyl]-2-thiazolyl]-5-methoxy-2-methyl- (CA INDEX NAME)



RN 773898-15-0 HCPLUS

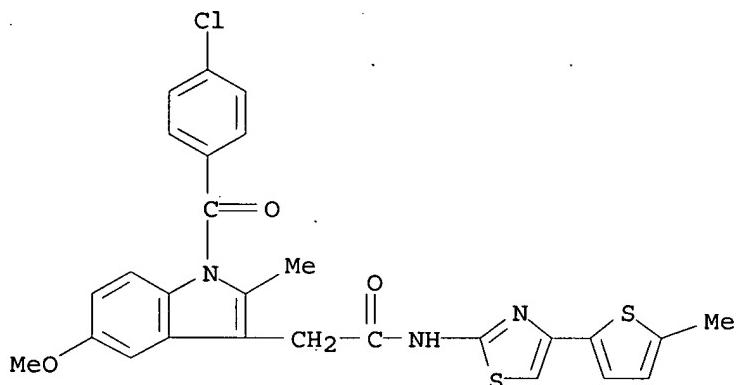
10542169.trn

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-[4-(3,4-dimethylphenyl)-2-thiazolyl]-5-methoxy-2-methyl- (CA INDEX NAME)



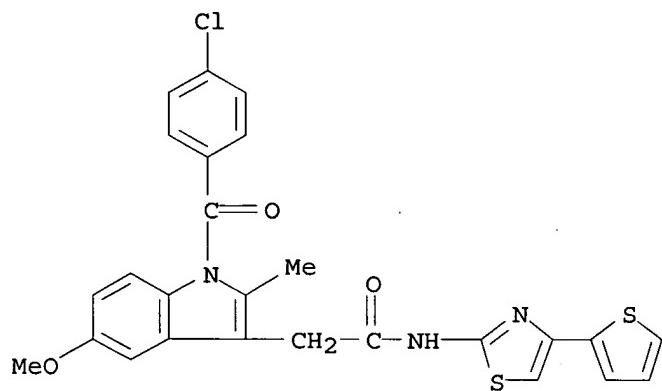
RN 773898-16-1 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[4-(5-methyl-2-thienyl)-2-thiazolyl]- (CA INDEX NAME)



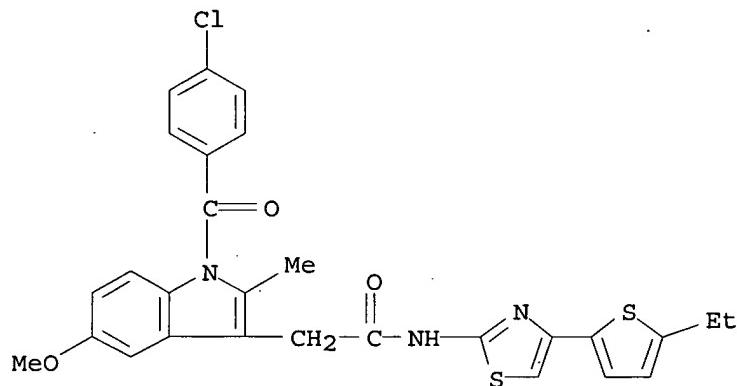
RN 773898-17-2 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[4-(2-thienyl)-2-thiazolyl]- (CA INDEX NAME)



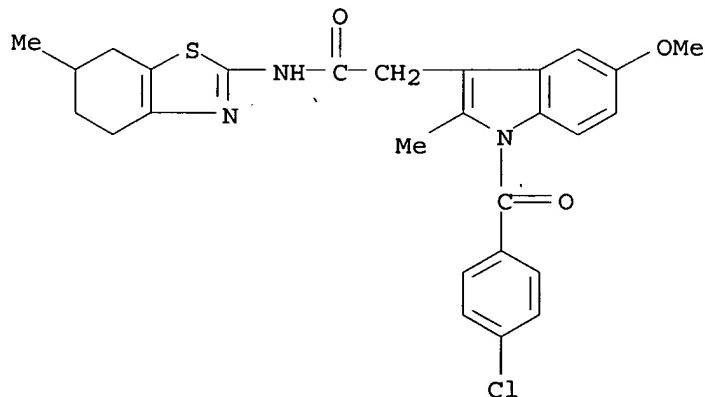
RN 773898-18-3 HCPLUS

CN 1H-Indole-3-acetamide, 1- (4-chlorobenzoyl)-N- [4- (5-ethyl-2-thienyl)-2-thiazolyl]-5-methoxy-2-methyl- (CA INDEX NAME)



RN 773898-19-4 HCPLUS

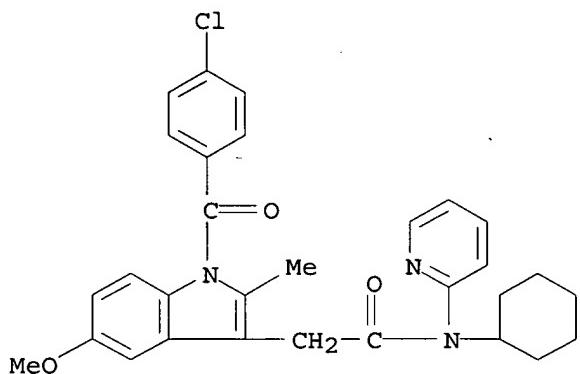
CN 1H-Indole-3-acetamide, 1- (4-chlorobenzoyl)-5-methoxy-2-methyl-N- (4,5,6,7-tetrahydro-6-methyl-2-benzothiazolyl)- (CA INDEX NAME)



RN 773898-32-1 HCPLUS

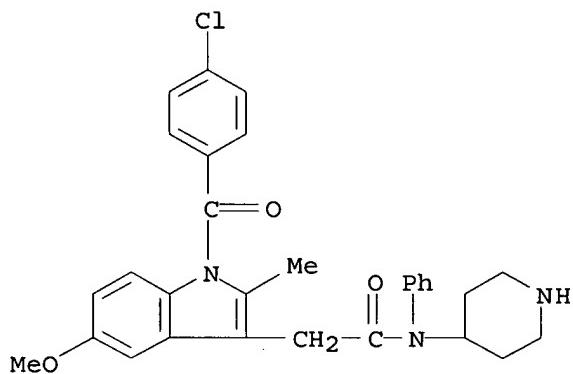
10542169.trn

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-2-pyridinyl- (CA INDEX NAME)



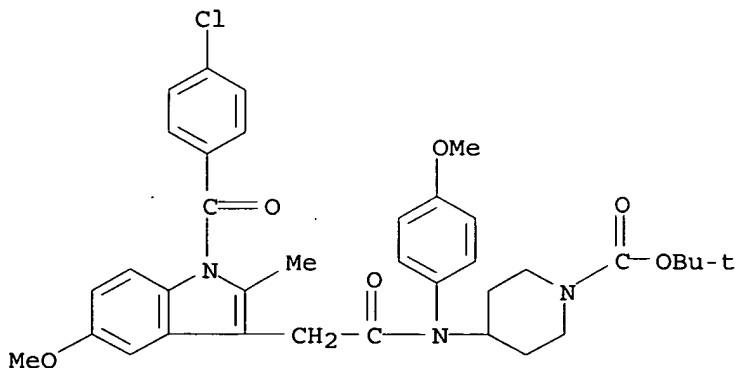
RN 773898-33-2 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-phenyl-N-4-piperidinyl- (CA INDEX NAME)



RN 773898-35-4 HCPLUS

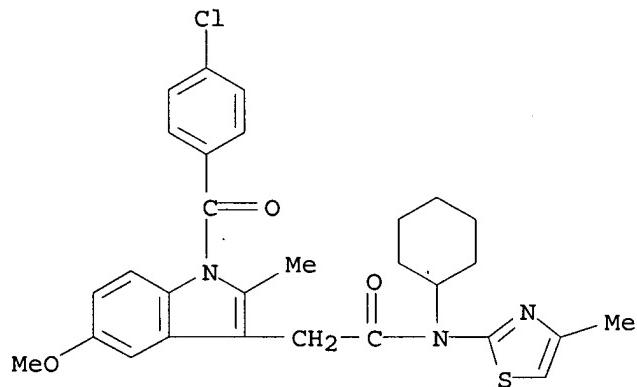
CN 1-Piperidinecarboxylic acid, 4-[[[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]acetyl](4-methoxyphenyl)amino]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



10542169.trn

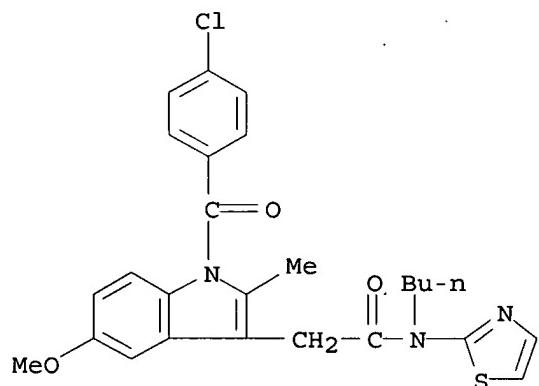
RN 773898-36-5 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-(4-methyl-2-thiazolyl)- (CA INDEX NAME)



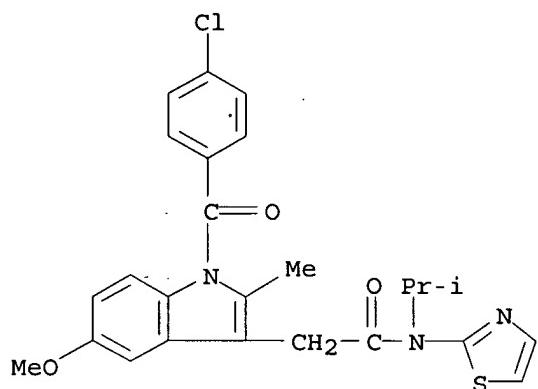
RN 773898-37-6 HCPLUS

CN 1H-Indole-3-acetamide, N-butyl-1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



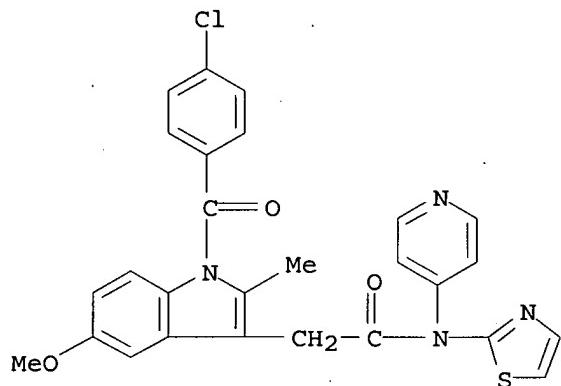
RN 773898-38-7 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(1-methylethyl)-N-2-thiazolyl- (CA INDEX NAME)



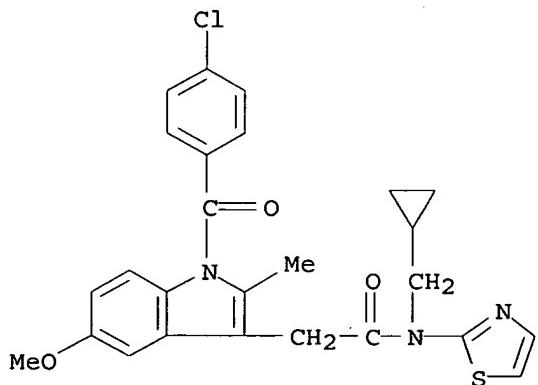
RN 773898-39-8 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-4-pyridinyl-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-40-1 HCPLUS

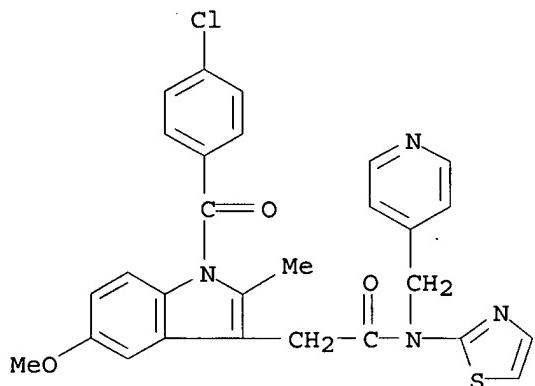
CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-41-2 HCPLUS

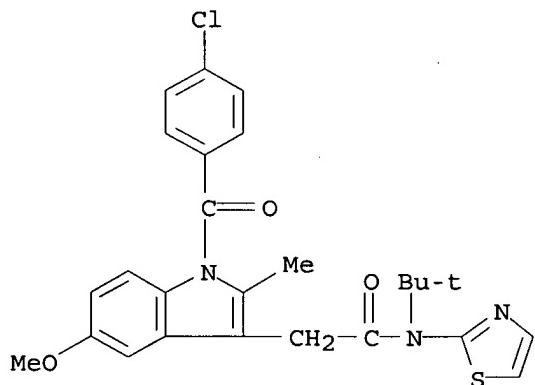
10542169.trn

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(4-pyridinylmethyl)-N-2-thiazolyl- (CA INDEX NAME)



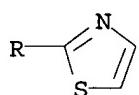
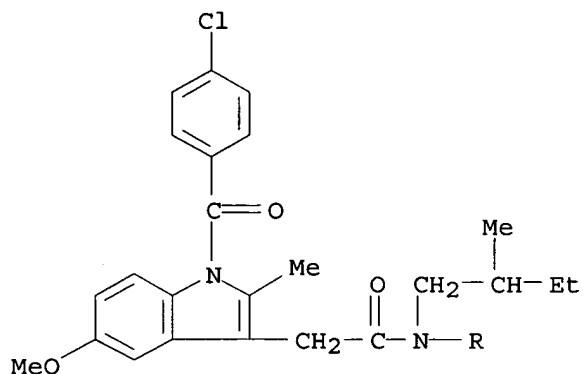
RN 773898-42-3 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-(1,1-dimethylethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



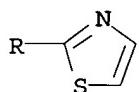
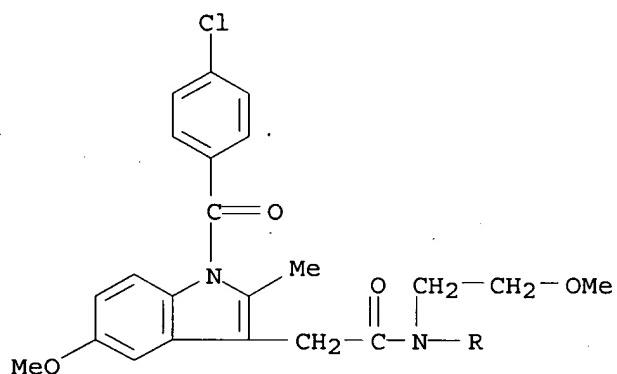
RN 773898-43-4 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(2-methylbutyl)-N-2-thiazolyl- (CA INDEX NAME)



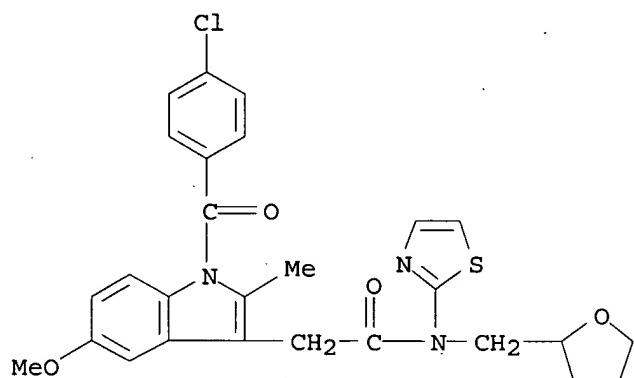
RN 773898-44-5 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-N-(2-methoxyethyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



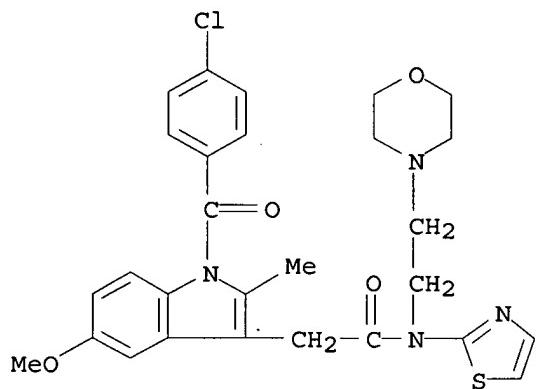
RN 773898-45-6 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[(tetrahydro-2-furanyl)methyl]-N-2-thiazolyl- (CA INDEX NAME)



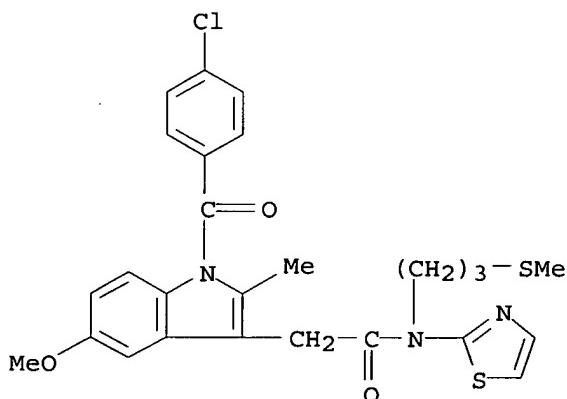
RN 773898-46-7 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[2-(4-morpholinyl)ethyl]-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-47-8 HCAPLUS

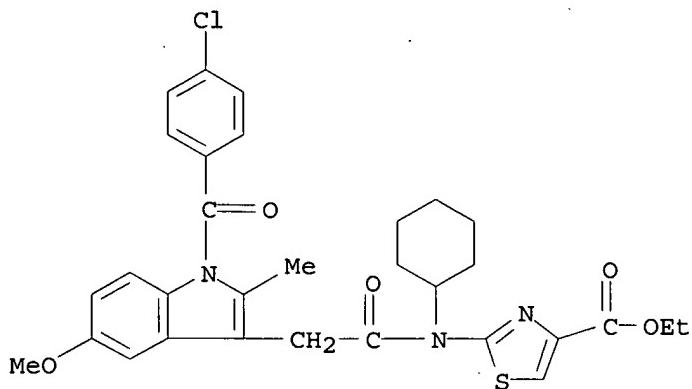
CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[3-(methylthio)propyl]-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-48-9 HCAPLUS

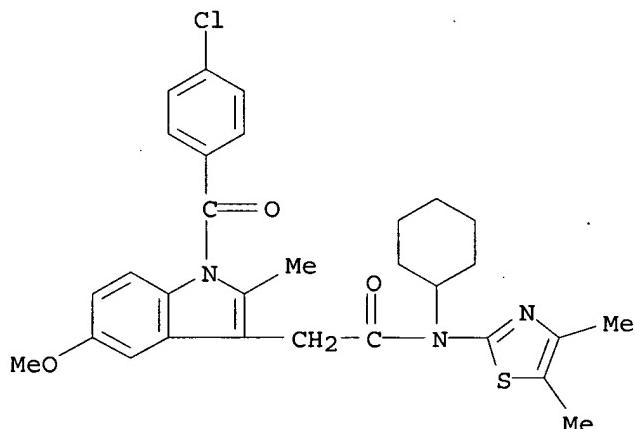
CN 4-Thiazolecarboxylic acid, 2-[[[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-

indol-3-yl]acetyl]cyclohexylamino]-, ethyl ester (9CI) (CA INDEX NAME)



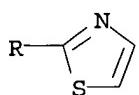
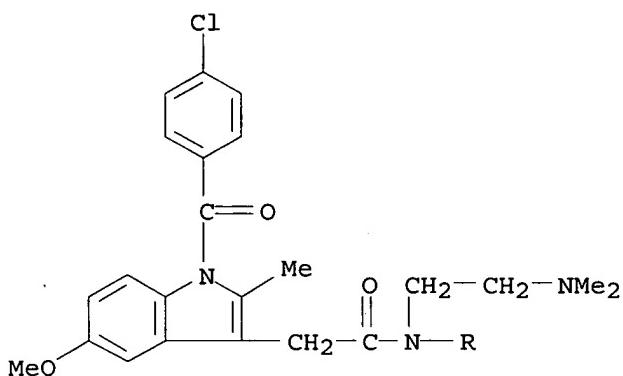
RN 773898-49-0 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-N-(4,5-dimethyl-2-thiazolyl)-5-methoxy-2-methyl- (CA INDEX NAME)



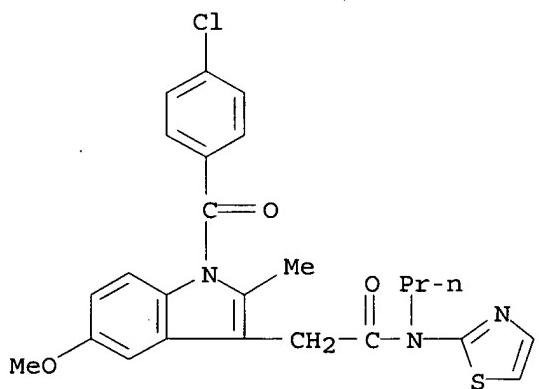
RN 773898-50-3 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-[2-(dimethylamino)ethyl]-5-methoxy-2-methyl-N-(2-thiazolyl)- (CA INDEX NAME)



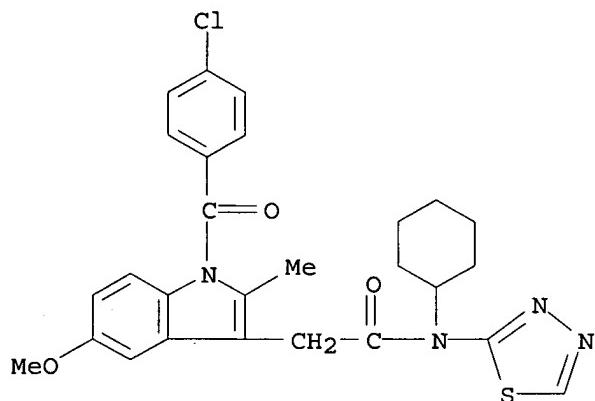
RN 773898-51-4 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-propyl-N-2-thiazolyl- (CA INDEX NAME)



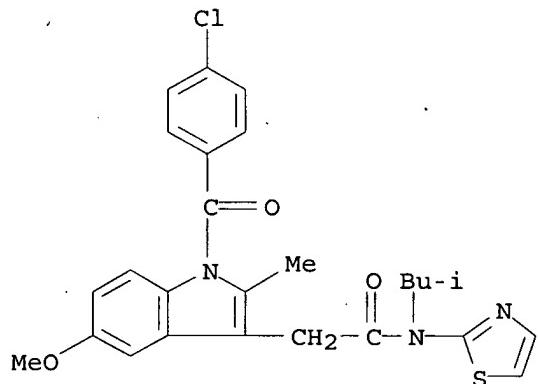
RN 773898-52-5 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-1,3,4-thiadiazol-2-yl- (CA INDEX NAME)



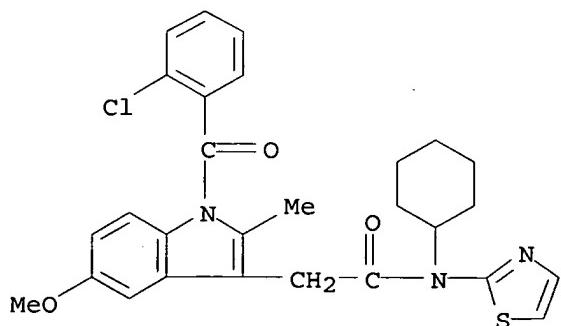
RN 773898-53-6 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(2-methylpropyl)-N-2-thiazolyl- (CA INDEX NAME)



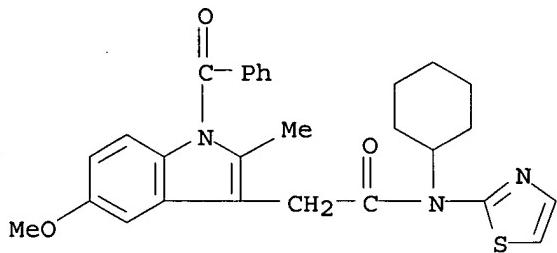
RN 773898-54-7 HCPLUS

CN 1H-Indole-3-acetamide, 1-(2-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



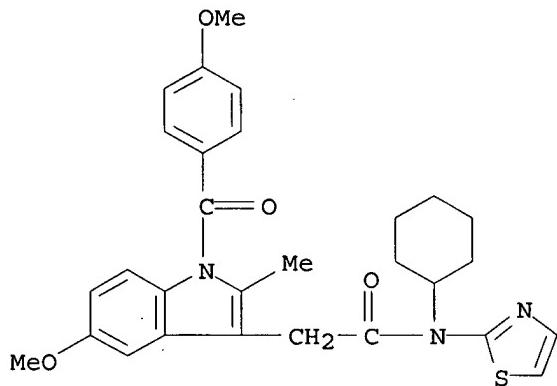
RN 773898-55-8 HCPLUS

CN 1H-Indole-3-acetamide, 1-benzoyl-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



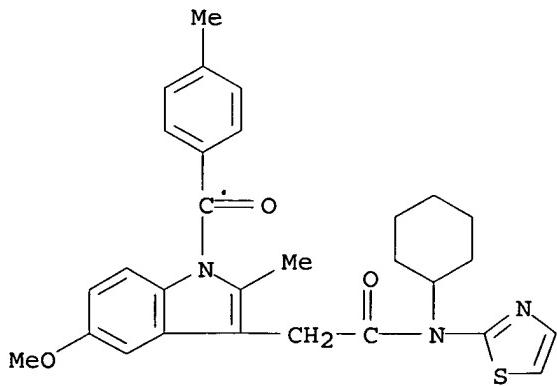
RN 773898-56-9 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-1-(4-methoxybenzoyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



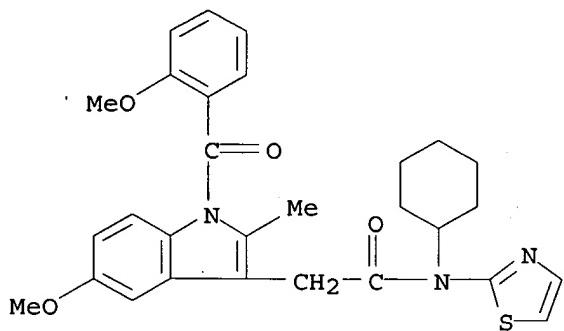
RN 773898-57-0 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-1-(4-methylbenzoyl)-N-2-thiazolyl- (CA INDEX NAME)

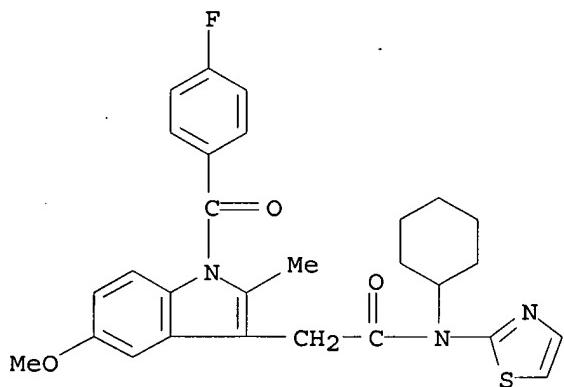


RN 773898-58-1 HCAPLUS

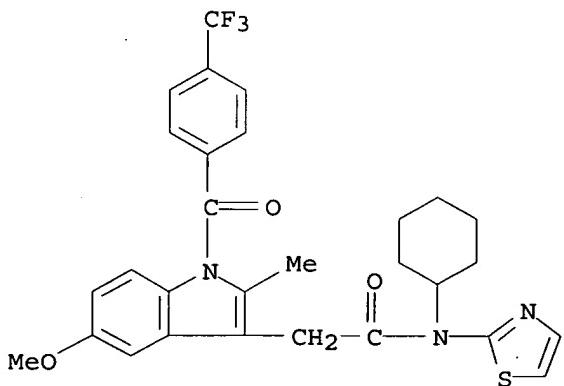
CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-1-(2-methoxybenzoyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



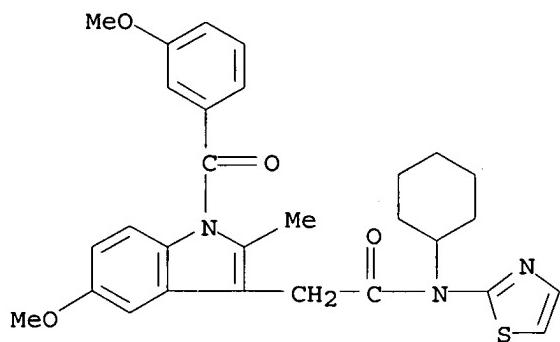
RN 773898-59-2 HCAPLUS
CN 1H-Indole-3-acetamide, N-cyclohexyl-1-(4-fluorobenzoyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



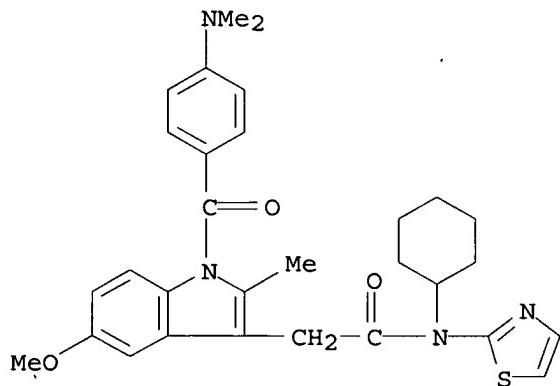
RN 773898-60-5 HCAPLUS
CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-[4-(trifluoromethyl)benzoyl]- (CA INDEX NAME)



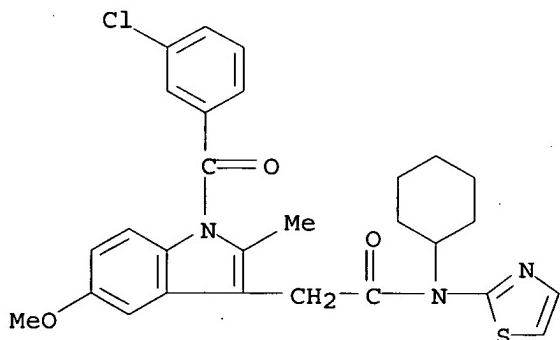
RN 773898-61-6 HCAPLUS
CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-1-(3-methoxybenzoyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



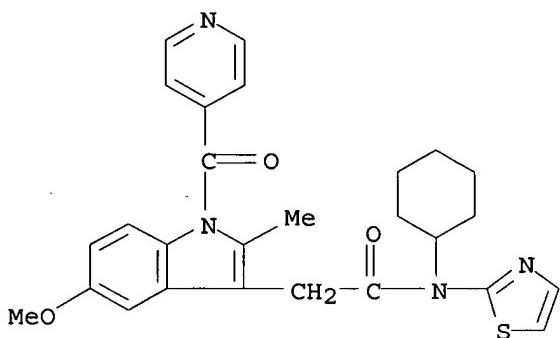
RN 773898-62-7 HCAPLUS
CN 1H-Indole-3-acetamide, N-cyclohexyl-1-[4-(dimethylamino)benzoyl]-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-63-8 HCAPLUS
CN 1H-Indole-3-acetamide, 1-(3-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)

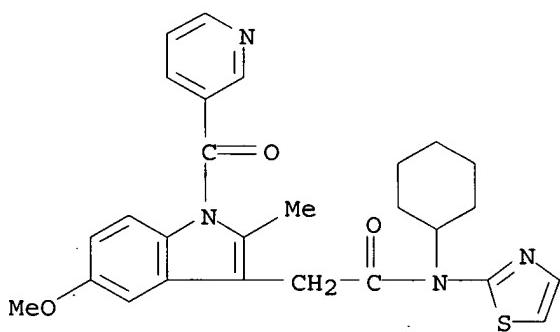


RN 773898-65-0 HCAPLUS
CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-1-(4-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



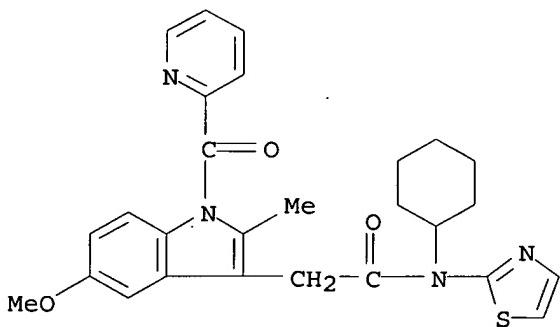
RN 773898-66-1 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-1-(3-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



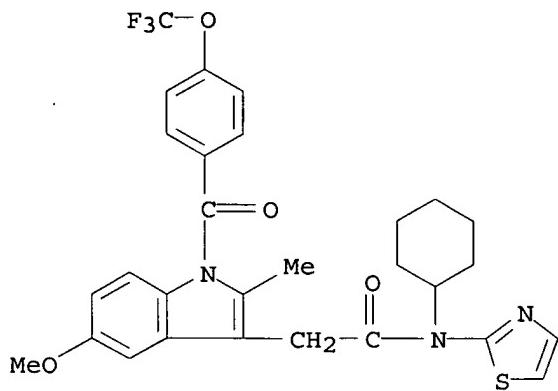
RN 773898-67-2 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-1-(2-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



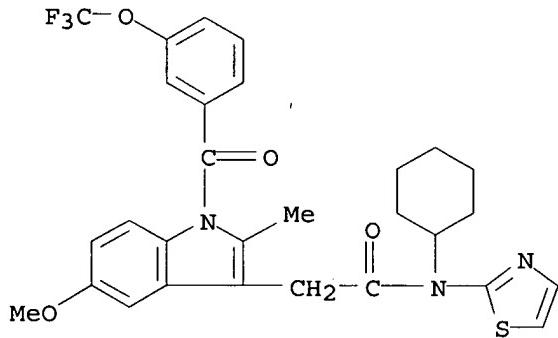
RN 773898-69-4 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-[4-(trifluoromethoxy)benzoyl]- (CA INDEX NAME)



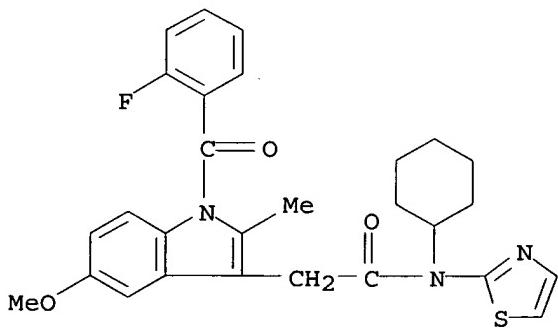
RN 773898-70-7 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-[3-(trifluoromethoxy)benzoyl]- (CA INDEX NAME)



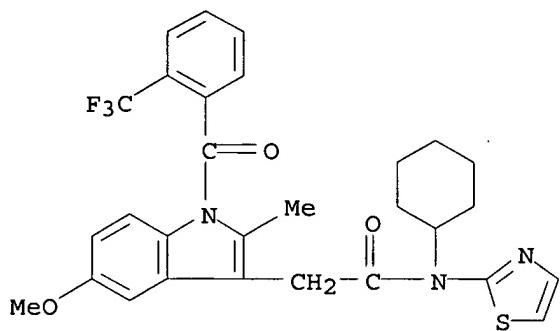
RN 773898-71-8 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-1-(2-fluorobenzoyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



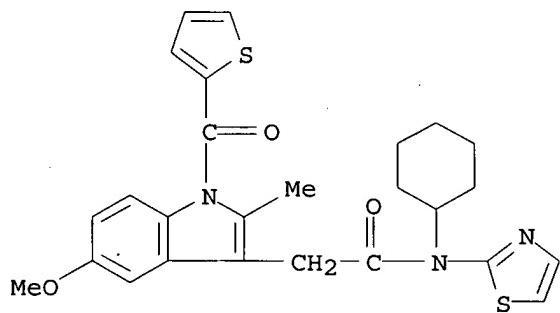
RN 773898-72-9 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-[2-(trifluoromethyl)benzoyl]- (CA INDEX NAME)



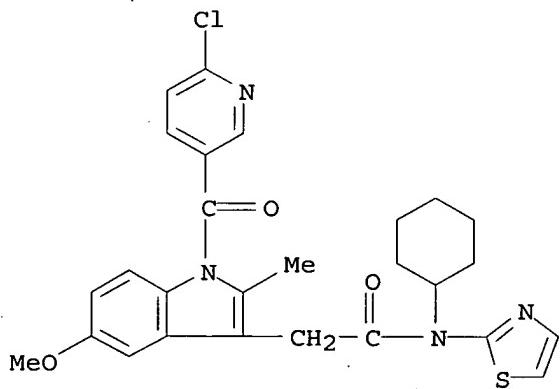
RN 773898-73-0 HCPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-(2-thienylcarbonyl)- (CA INDEX NAME)



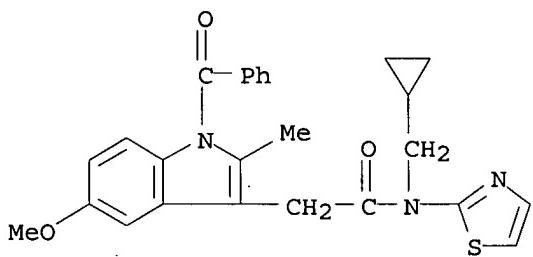
RN 773898-75-2 HCPLUS

CN 1H-Indole-3-acetamide, 1-[(6-chloro-3-pyridinyl)carbonyl]-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



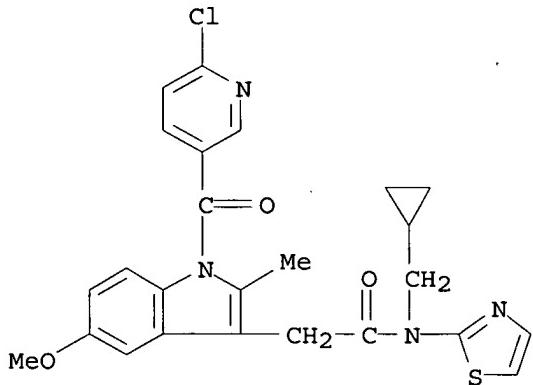
RN 773898-77-4 HCPLUS

CN 1H-Indole-3-acetamide, 1-benzoyl-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



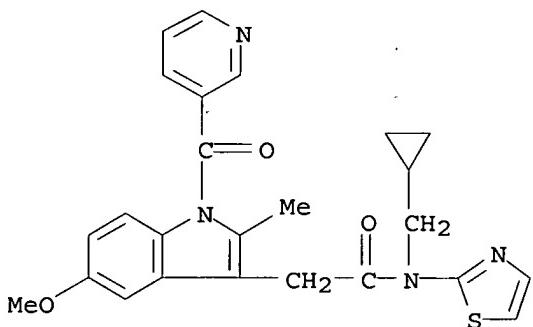
RN 773898-78-5 HCAPLUS

CN 1H-Indole-3-acetamide, 1-[(6-chloro-3-pyridinyl)carbonyl]-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



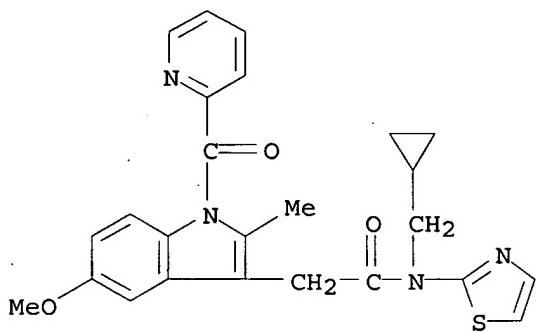
RN 773898-79-6 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(3-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



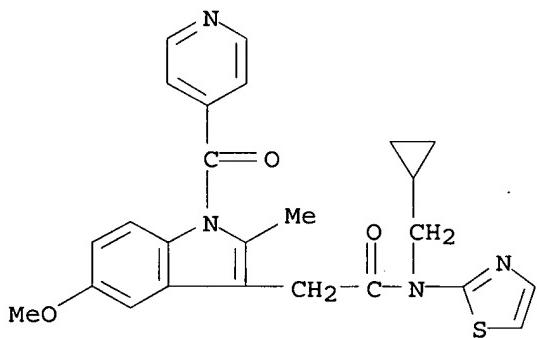
RN 773898-80-9 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(2-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



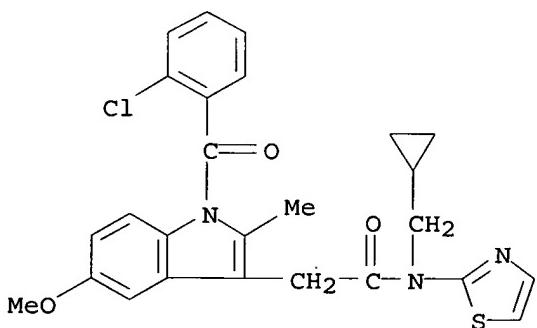
RN 773898-81-0 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(4-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



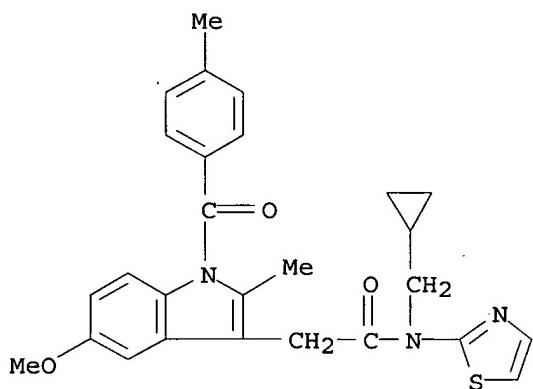
RN 773898-82-1 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(2-chlorobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



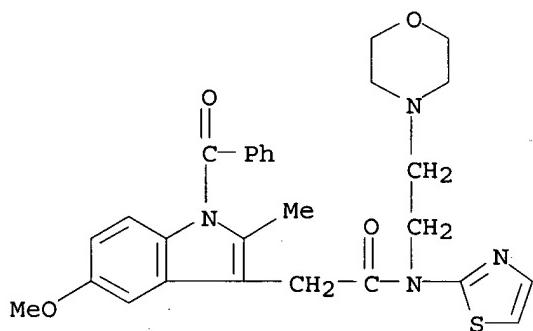
RN 773898-83-2 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(4-methylbenzoyl)-N-2-thiazolyl- (CA INDEX NAME)



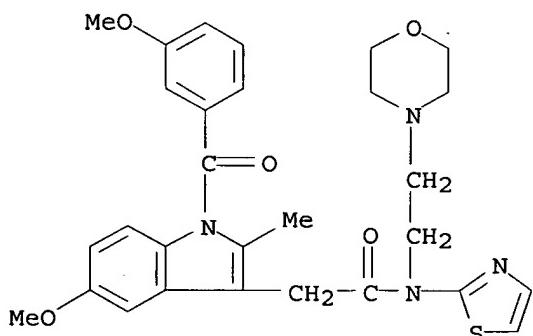
RN 773898-84-3 HCAPLUS

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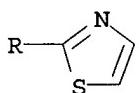
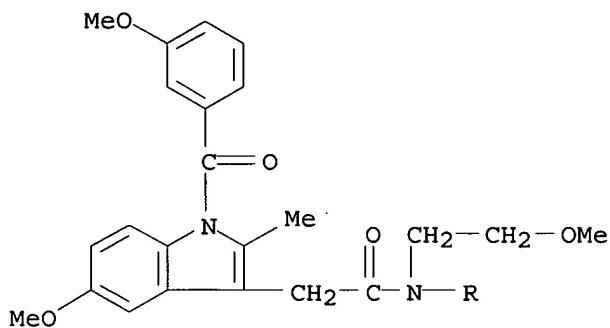
RN 773898-85-4 HCAPLUS

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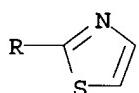
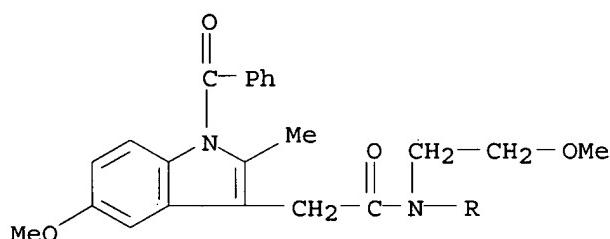
RN 773898-86-5 HCAPLUS

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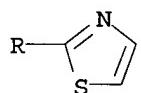
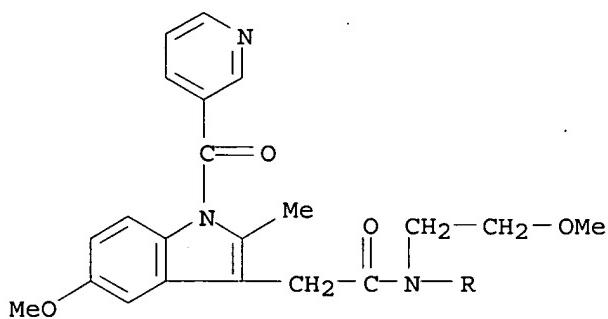
RN 773898-87-6 HCPLUS

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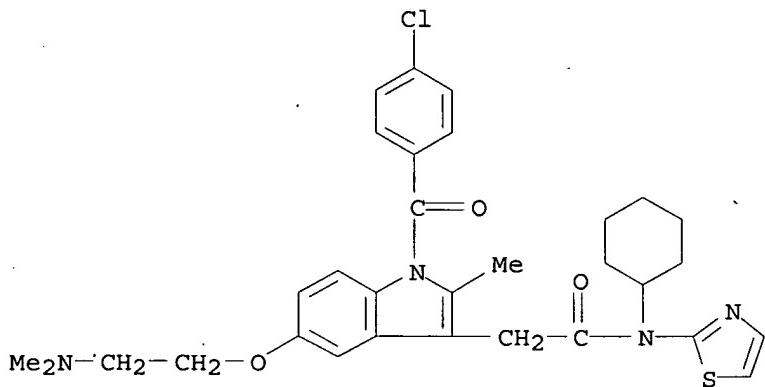
RN 773898-88-7 HCPLUS

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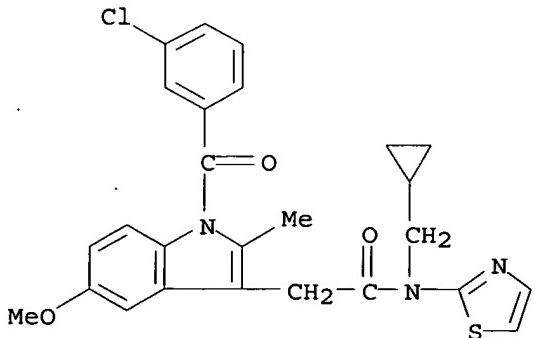
RN 773898-90-1 HCAPLUS

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RN 773898-91-2 HCAPLUS

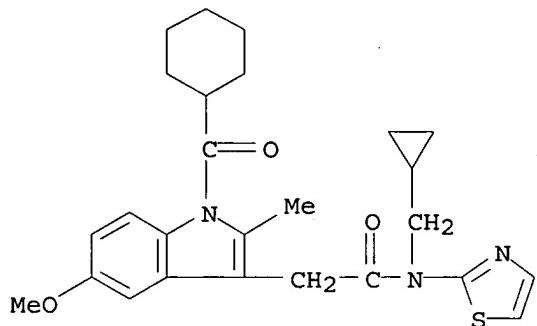
CN 1H-Indole-3-acetamide, 1-(3-chlorobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



10542169.trn

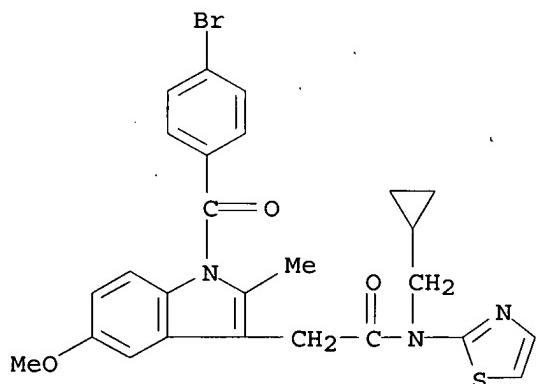
RN 773898-92-3 HCAPLUS

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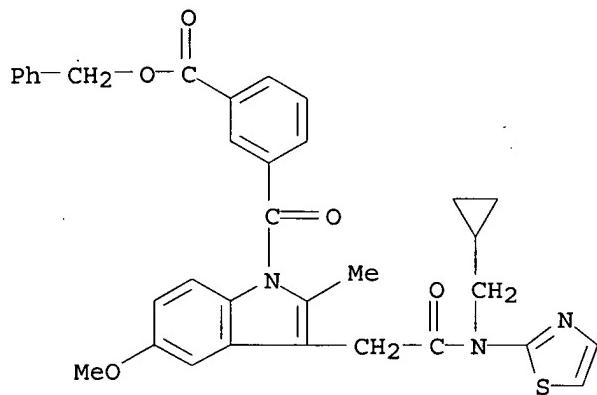
RN 773898-93-4 HCAPLUS

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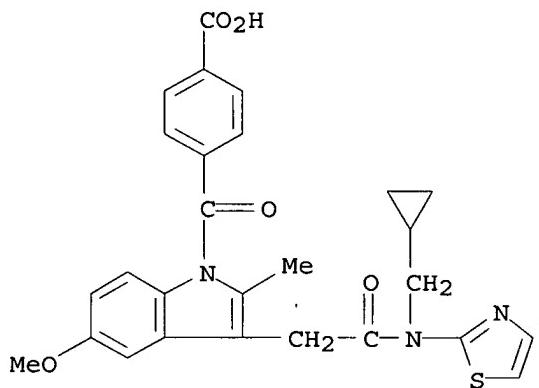
RN 773898-94-5 HCAPLUS

CN Benzoic acid, 3-[[3-[2-[(cyclopropylmethyl)-2-thiazolylamino]-2-oxoethyl]-5-methoxy-2-methyl-1H-indol-1-yl]carbonyl]-, phenylmethyl ester (CA INDEX NAME)



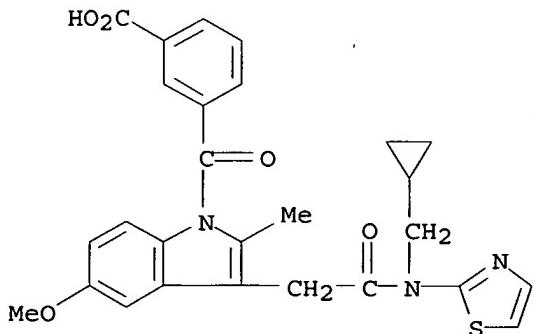
RN 773898-97-8 HCAPLUS

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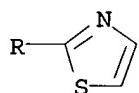
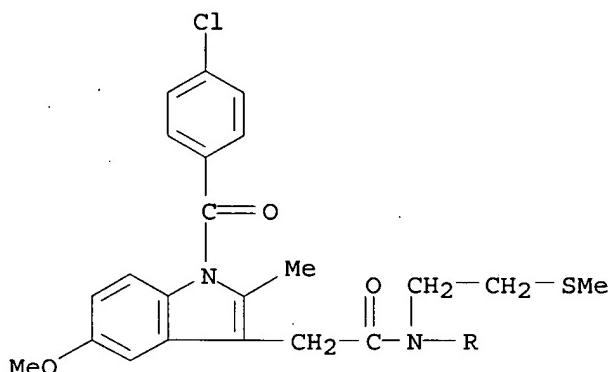
RN 773898-98-9 HCAPLUS

CN Benzoic acid, 3-[[3-[(cyclopropylmethyl)-2-thiazolylamino]-2-oxoethyl]-5-methoxy-2-methyl-1H-indol-1-yl]carbonyl] - (CA INDEX NAME)



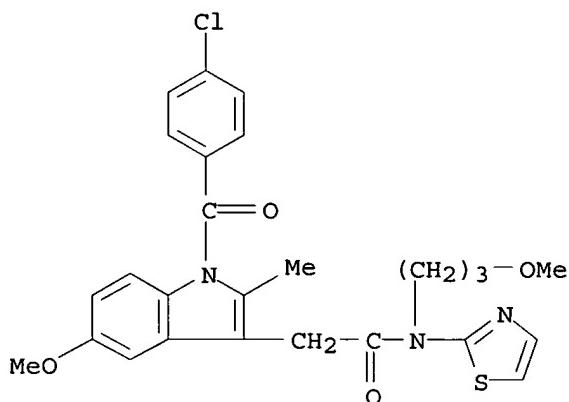
RN 773898-99-0 HCAPLUS

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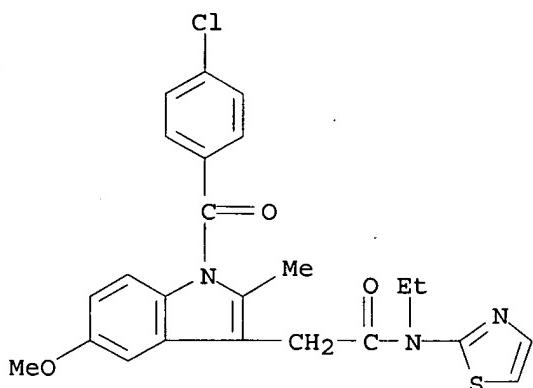
RN 773899-00-6 HCPLUS

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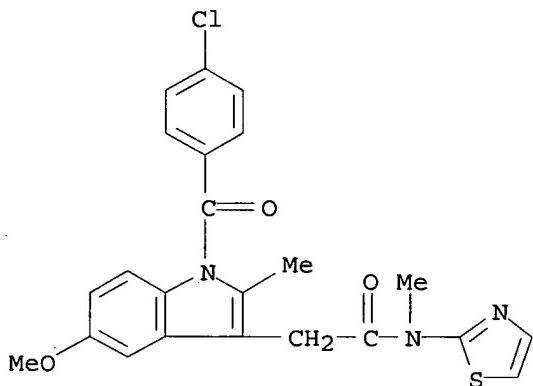
RN 773899-01-7 HCPLUS

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RN 773899-02-8 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-N,2-dimethyl-N-2-thiazolyl- (CA INDEX NAME)



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L18 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:857325 HCAPLUS

DOCUMENT NUMBER: 141:350033

TITLE: Preparation of 5-methoxy-2-methylindole-3-acetamide derivs. as potassium channel blockers for treating ocular hypertension

INVENTOR(S):

Fisher, Michael H.; Garcia, Maria L.; Kaczorowski,
Gregory J.; Meinke, Peter T.; Parsons, William H.;
 Boyd, Edward Andrew; Price, Stephen; Stibbard, John

PATENT ASSIGNEE(S):

Merck & Co., Inc., USA; Evotec Oai

SOURCE:

PCT Int. Appl., 109 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.

KIND DATE

APPLICATION NO.

DATE

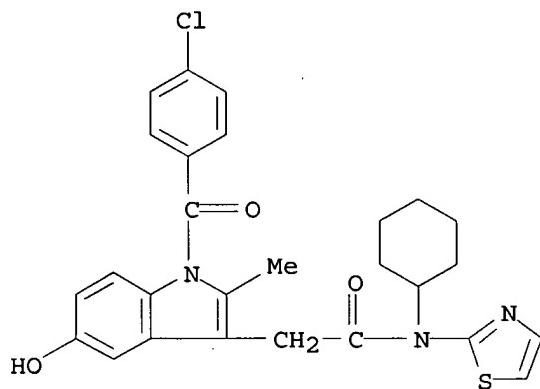
| WO 2004087051 | A2 | 20041014 | WO 2004-US9028 | 20040324 |
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| WO 2004087051 | A3 | 20050721 | | |
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| CN 1791402 | A | 20060621 | CN 2004-80013916 | 20040324 |
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| US 2006069256 | A1 | 20060330 | US 2005-542169 | 20050713 |
| IN 2005DN04100 | A | 20070831 | IN 2005-DN4100 | 20050912 |
| PRIORITY APPLN. INFO.: | | | US 2003-458103P | P 20030327 |
| | | | WO 2004-US9028 | A 20040324 |
| OTHER SOURCE(S): | CASREACT 141:350033; MARPAT 141:350033 | | | |
| GI | | | | |

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The title compds. I [X = -(CHR7)p-; Y = -CO(CH2)n- or -CH(OR8)-; Q = N, CR9, or O; R1 = H, alkyl, CF3, alkoxy, OH, etc.; R2 = H, alkyl, alkylSR8, -(CH2)nO(CH2)mOR8, -(CH2)alkoxy, etc.; R3 = H, alkyl, -(CH2)ncycloalkyl, -(CH2)nheterocyclyl, or when Q = N, R2, R3 taken together with the the N form a 4-10 membered heterocyclic ring; R4, R5 = H, alkoxy, OH, alkyl, COOR8, SO3H, etc.; R6 = H, alkyl, -(CH2)(hetero)aryl, -NH(CH2)(hetero)aryl, etc.; R7 = H, alkyl, -(CH2)nCOOR8, or -(CH2)nN(R8)2; R8 = H, or alkyl; R9 = H, or alkyl; m = 0-3; n = 0-3, p = 0-1] were prepared as potent potassium channel blockers in the treatment of glaucoma and other conditions which leads to elevated intraocular pressure in the eye of a patient. For example, reaction of 1-(4-chlorobenzoyl)-5-methoxy-2-methylindole-3-acetic acid with N-cyclohexyl-N-thiazol-2-yl amine (preparation given) yielded compound II. The compds. of this invention inhibited Maxi-K Channel activity with IC50's in the range of 1 nM to 20 μ M.

IT 773898-89-8P
 RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
 (preparation of 5-methoxy-2-methylindole-3-acetamide derivs. as potassium channel blockers for treating ocular hypertension)

RN 773898-89-8 HCAPLUS
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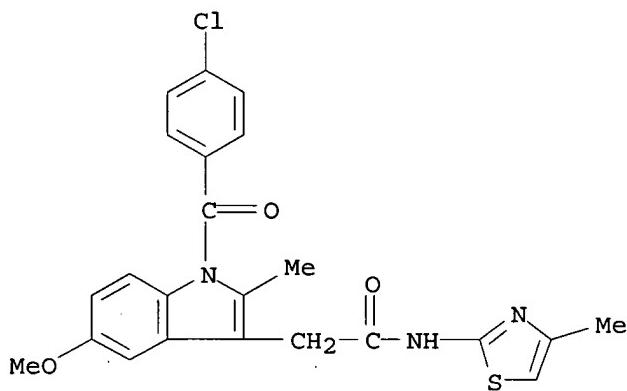
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 773898-99-0P 773899-00-6P 773899-01-7P
 773899-02-8P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of 5-methoxy-2-methylindole-3-acetamide derivs. as potassium channel blockers for treating ocular hypertension)

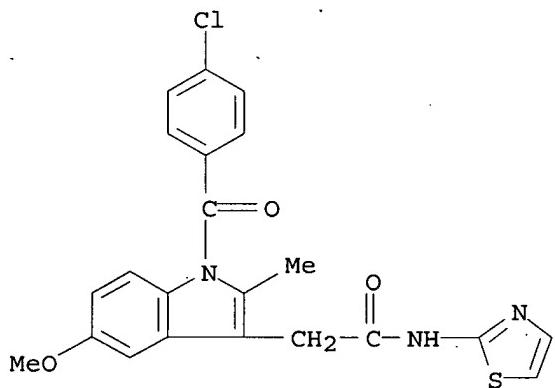
RN 282728-83-0 HCAPLUS

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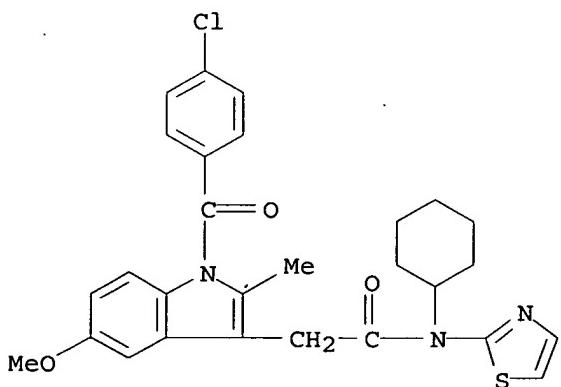
RN 732255-43-5 HCAPLUS

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RN 773898-05-8 HCAPLUS

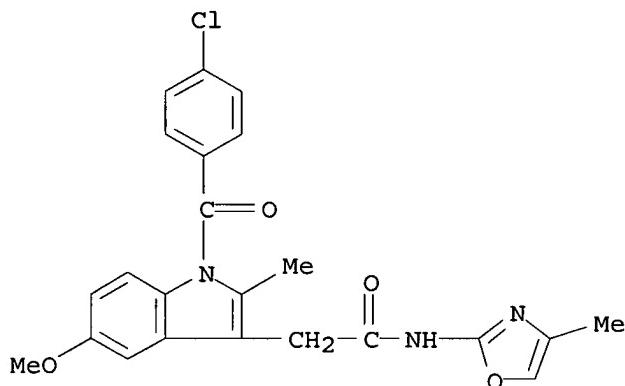
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RN 773898-10-5 HCAPLUS

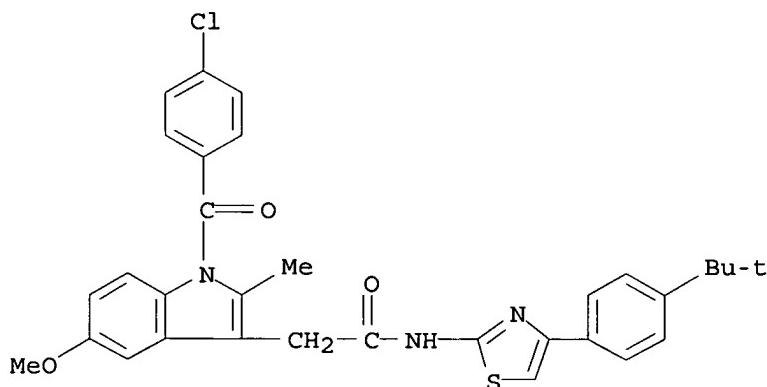
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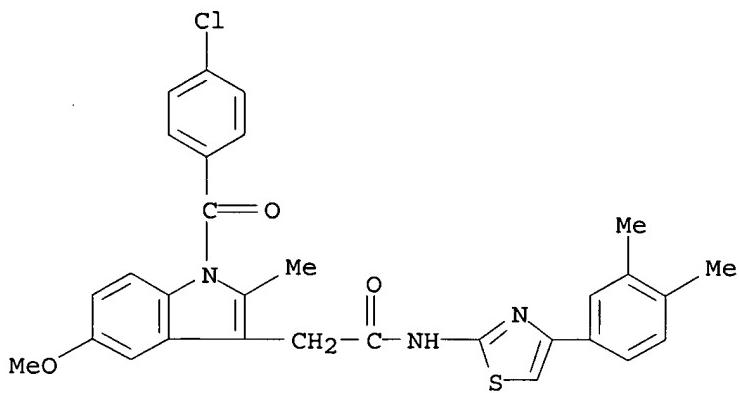
RN 773898-14-9 HCPLUS

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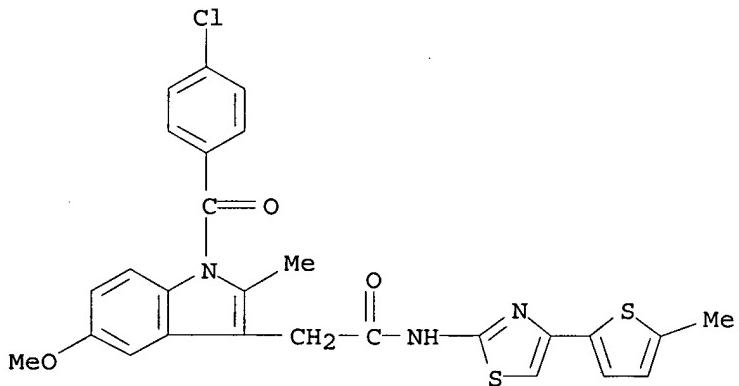
RN 773898-15-0 HCPLUS

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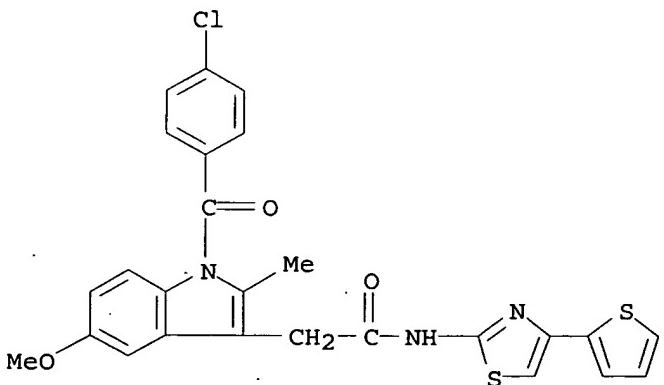
RN 773898-16-1 HCAPLUS

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RN 773898-17-2 HCAPLUS

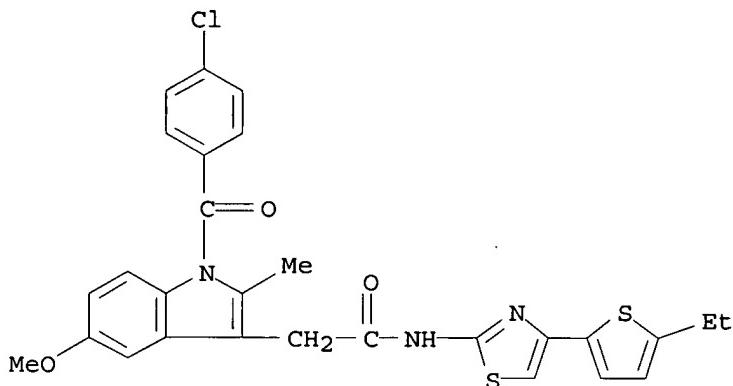
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RN 773898-18-3 HCAPLUS

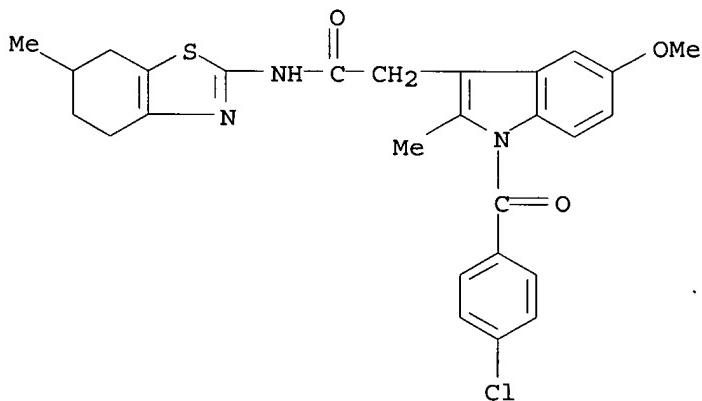
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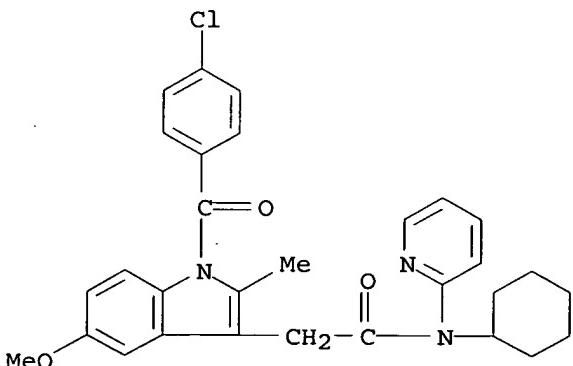
RN 773898-19-4 HCAPLUS

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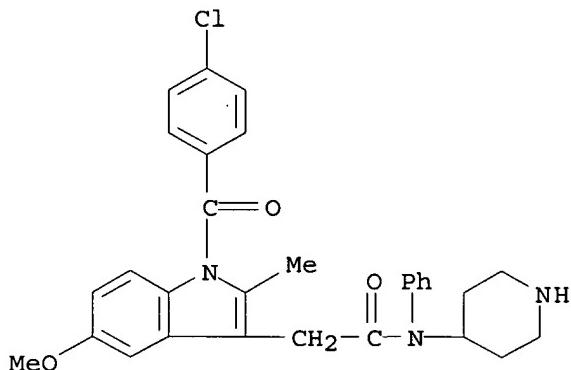


RN 773898-32-1 HCAPLUS

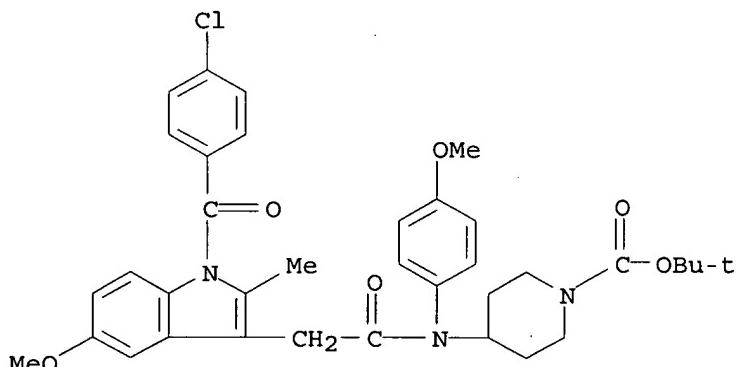
CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-2-pyridinyl- (CA INDEX NAME)



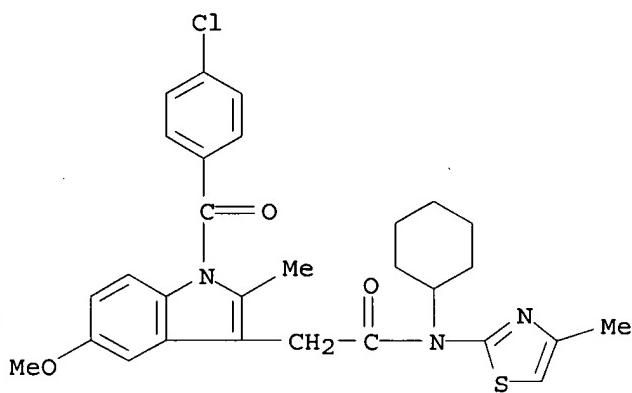
RN 773898-33-2 HCAPLUS
CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-phenyl-N-4-piperidinyl- (CA INDEX NAME)



RN 773898-35-4 HCAPLUS
CN 1-Piperidinecarboxylic acid, 4-[[[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]acetyl](4-methoxyphenyl)amino]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

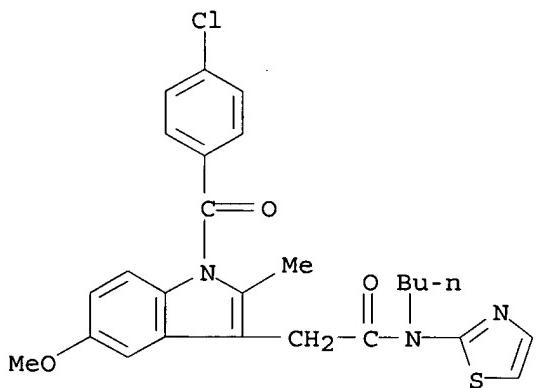


RN 773898-36-5 HCAPLUS
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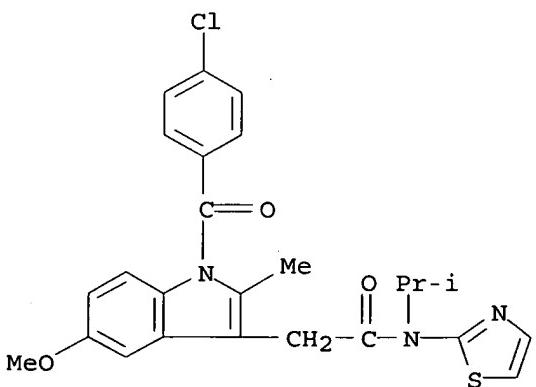
RN 773898-37-6 HCPLUS

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RN 773898-38-7 HCPLUS

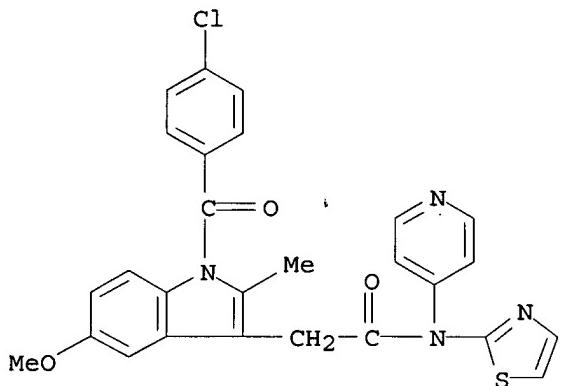
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RN 773898-39-8 HCPLUS

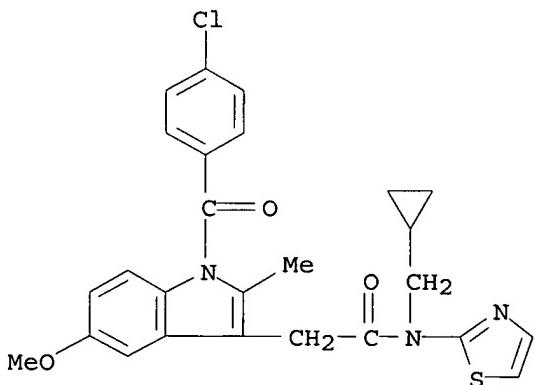
10542169.trn

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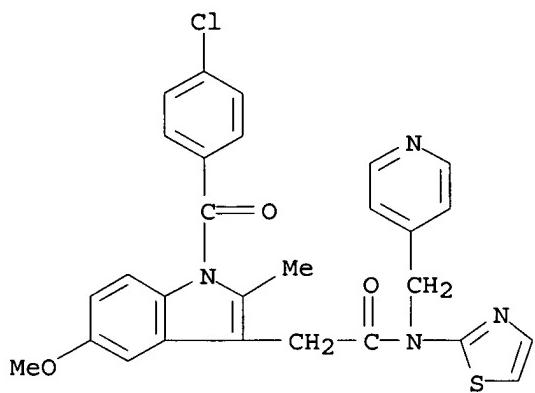
RN 773898-40-1 HCAPLUS

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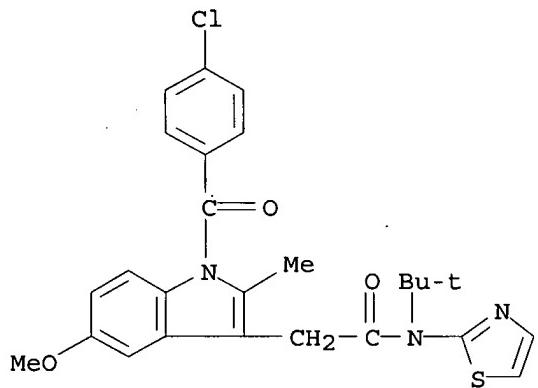
RN 773898-41-2 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(4-pyridinylmethyl)-N-2-thiazolyl- (CA INDEX NAME)



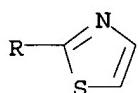
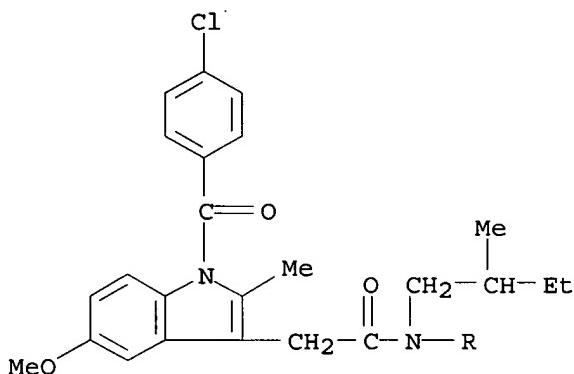
RN 773898-42-3 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-(1,1-dimethylethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



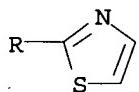
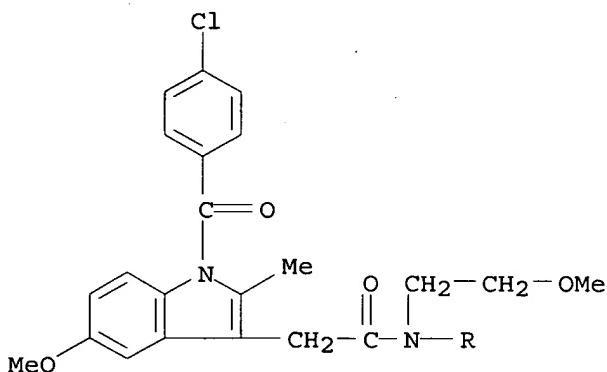
RN 773898-43-4 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(2-methylbutyl)-N-2-thiazolyl- (CA INDEX NAME)



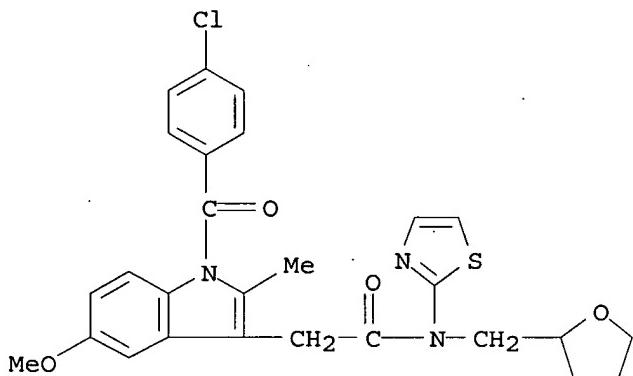
RN 773898-44-5 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-N-(2-methoxyethyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



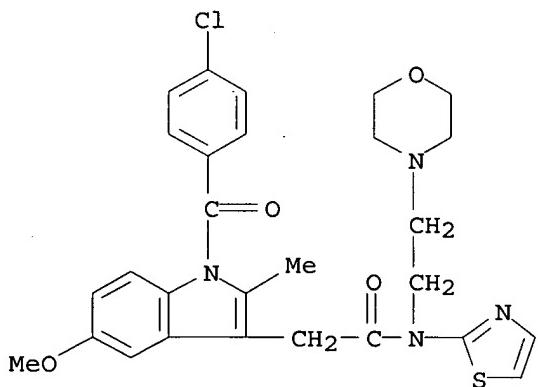
RN 773898-45-6 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[tetrahydro-2-furanyl]methyl]-N-2-thiazolyl- (CA INDEX NAME)



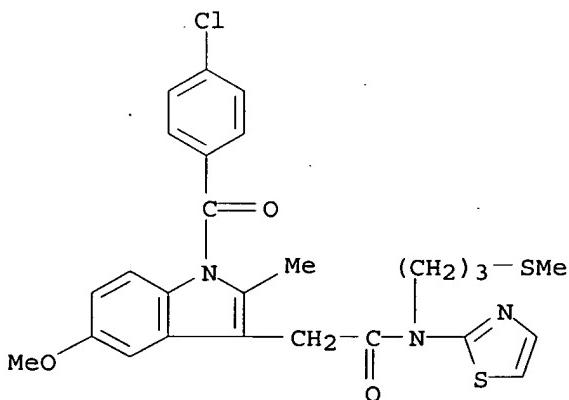
RN 773898-46-7 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[2-(4-morpholinyl)ethyl]-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-47-8 HCAPLUS

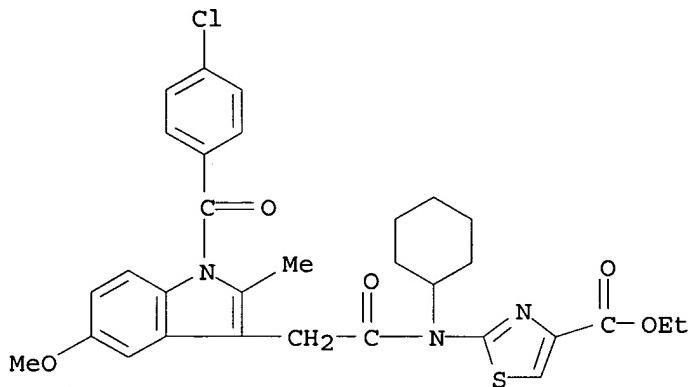
CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[3-(methylthio)propyl]-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-48-9 HCAPLUS

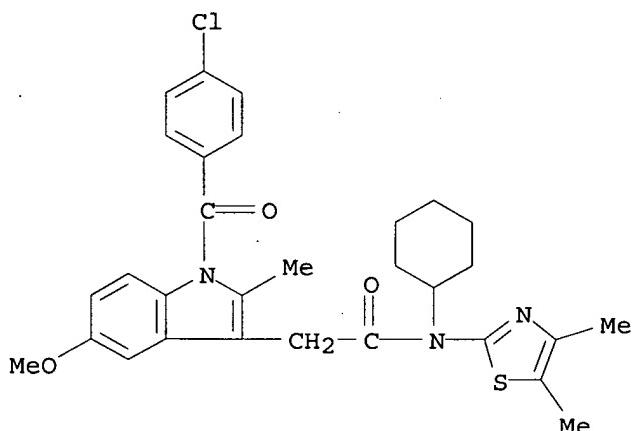
CN 4-Thiazolecarboxylic acid, 2-[[[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-

indol-3-yl]acetyl]cyclohexylamino]-, ethyl ester (9CI) (CA INDEX NAME)



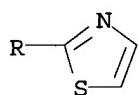
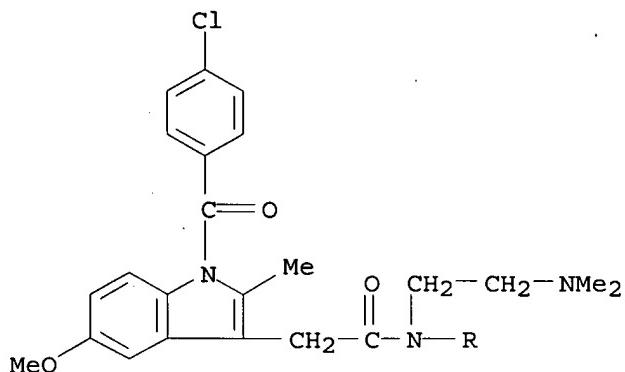
RN 773898-49-0 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-N-(4,5-dimethyl-2-thiazolyl)-5-methoxy-2-methyl- (CA INDEX NAME)



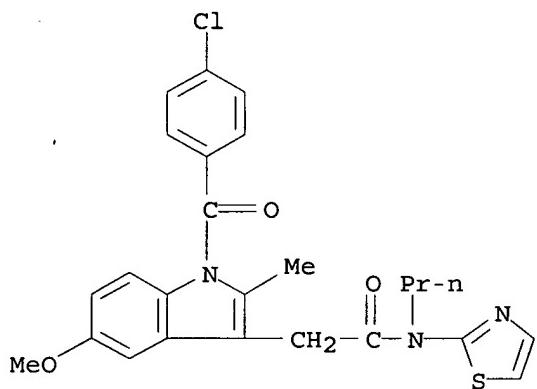
RN 773898-50-3 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-[2-(dimethylaminio)ethyl]-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



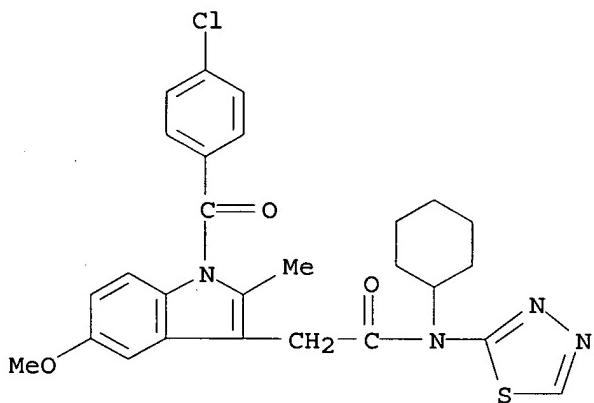
RN 773898-51-4 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-propyl-N-2-thiazolyl- (CA INDEX NAME)



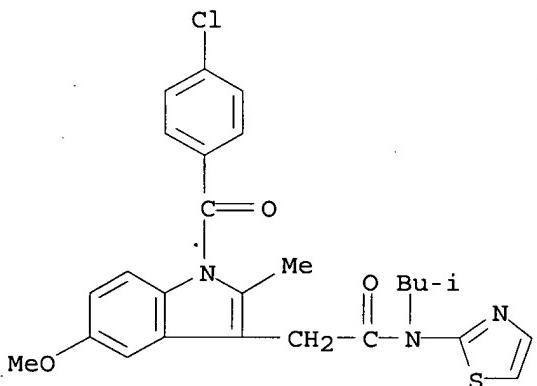
RN 773898-52-5 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-1,3,4-thiadiazol-2-yl- (CA INDEX NAME)



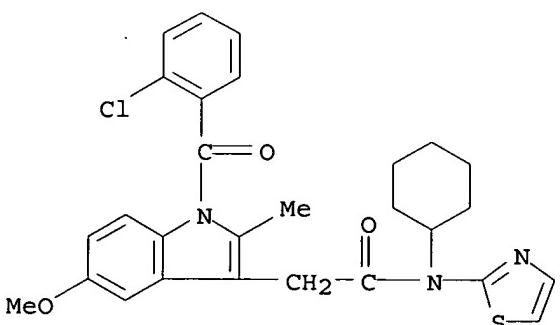
RN 773898-53-6 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-(2-methylpropyl)-N-2-thiazolyl- (CA INDEX NAME)



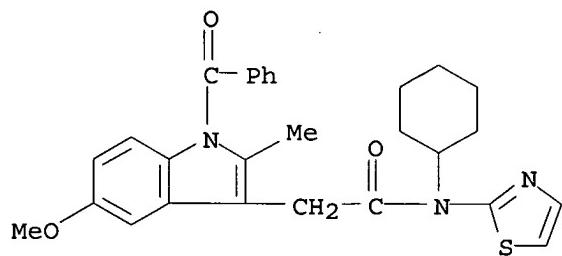
RN 773898-54-7 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(2-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



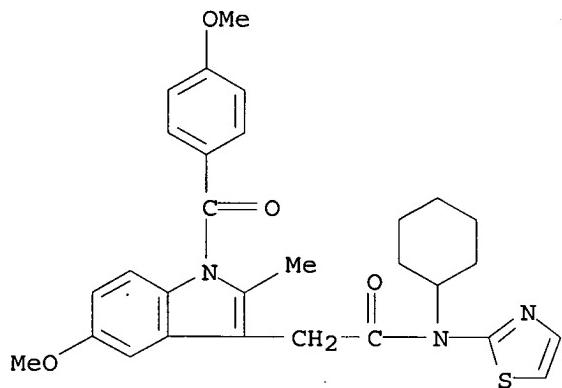
RN 773898-55-8 HCAPLUS

CN 1H-Indole-3-acetamide, 1-benzoyl-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



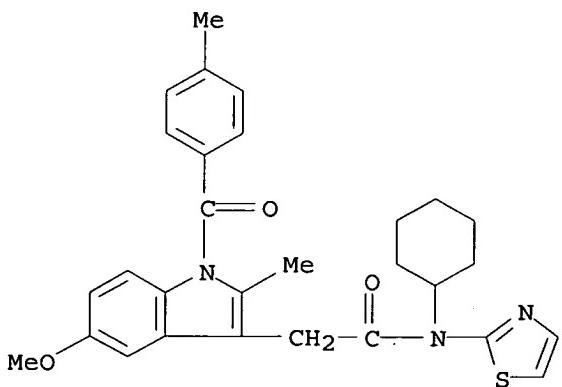
RN 773898-56-9 HCPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-1-(4-methoxybenzoyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



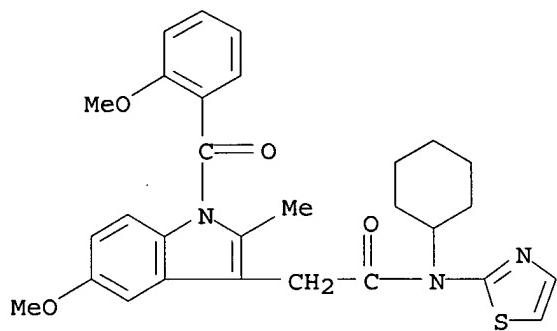
RN 773898-57-0 HCPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-1-(4-methylbenzoyl)-N-2-thiazolyl- (CA INDEX NAME)



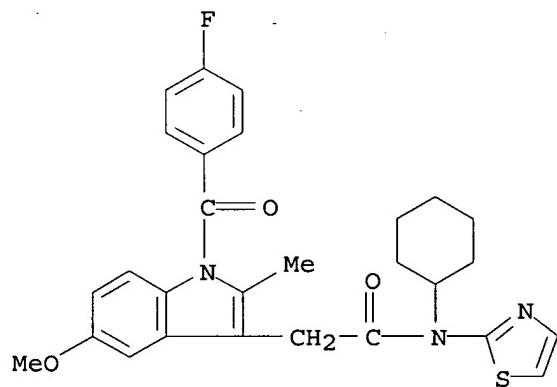
RN 773898-58-1 HCPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-1-(2-methoxybenzoyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



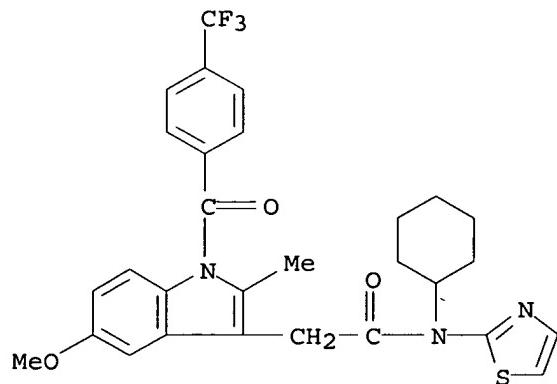
RN 773898-59-2 HCPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-1-(4-fluorobenzoyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



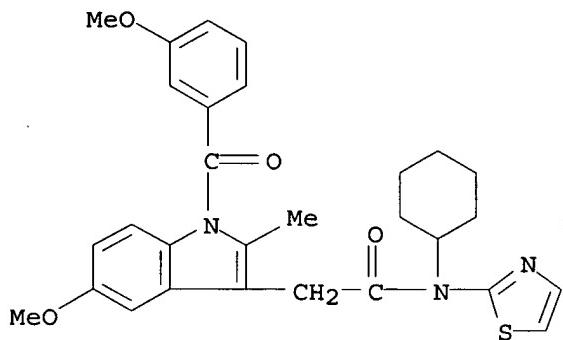
RN 773898-60-5 HCPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-[4-(trifluoromethyl)benzoyl]- (CA INDEX NAME)

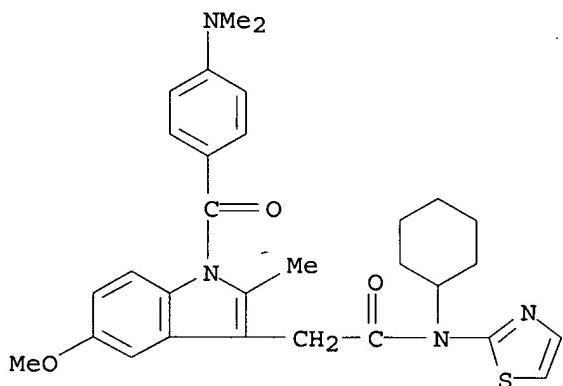


RN 773898-61-6 HCPLUS

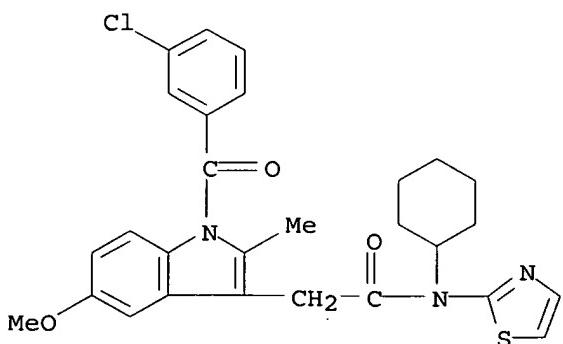
CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-1-(3-methoxybenzoyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



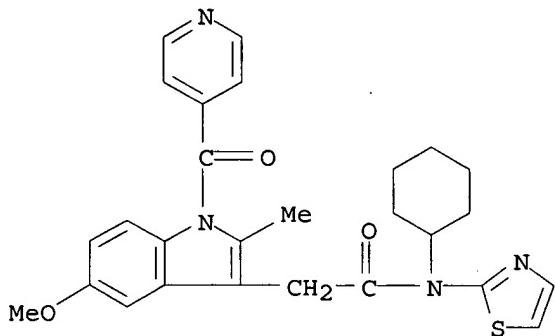
RN 773898-62-7 HCAPLUS
CN 1H-Indole-3-acetamide, N-cyclohexyl-1-[4-(dimethylamino)benzoyl]-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-63-8 HCAPLUS
CN 1H-Indole-3-acetamide, 1-(3-chlorobenzoyl)-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)

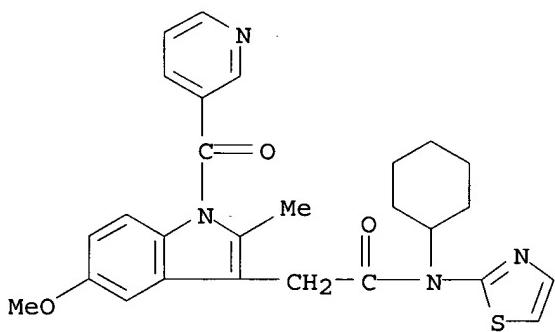


RN 773898-65-0 HCAPLUS
CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-1-(4-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



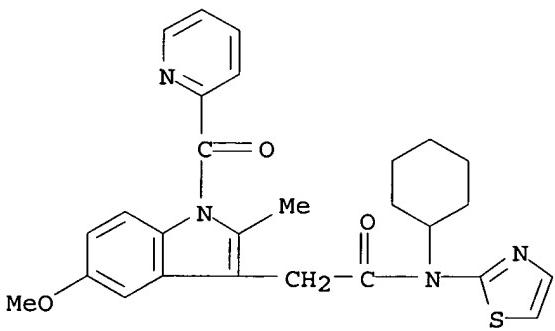
RN 773898-66-1 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-1-(3-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



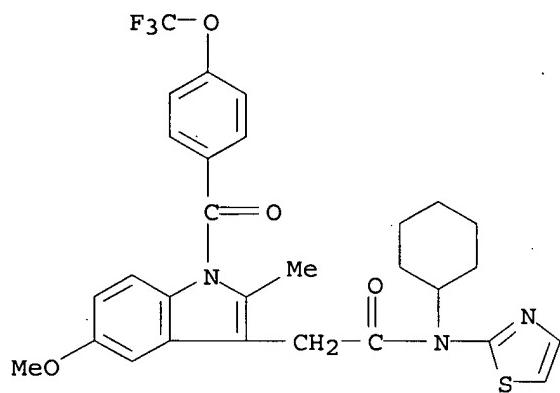
RN 773898-67-2 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-1-(2-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



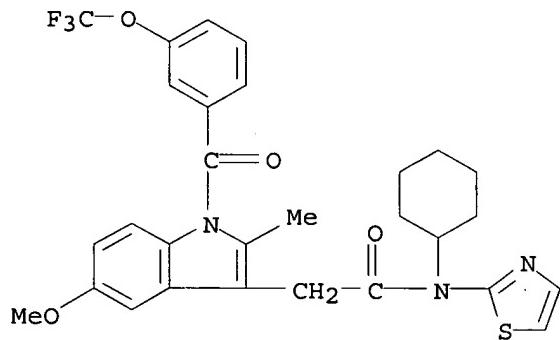
RN 773898-69-4 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-[4-(trifluoromethoxy)benzoyl]- (CA INDEX NAME)



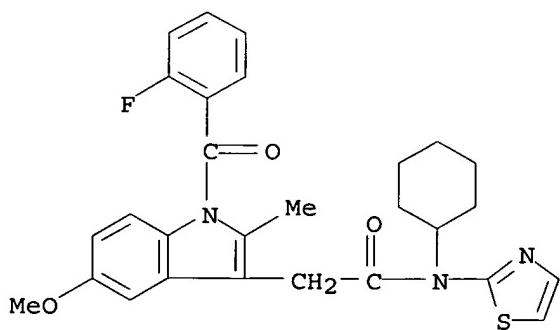
RN 773898-70-7 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-[3-(trifluoromethoxy)benzoyl]- (CA INDEX NAME)



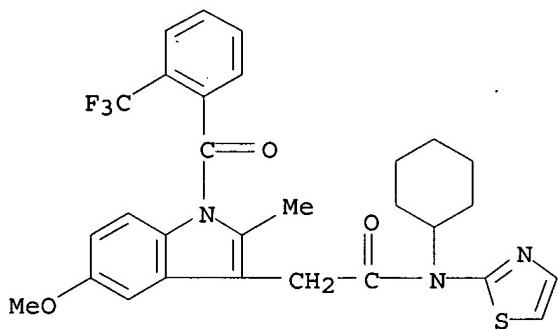
RN 773898-71-8 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-1-(2-fluorobenzoyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



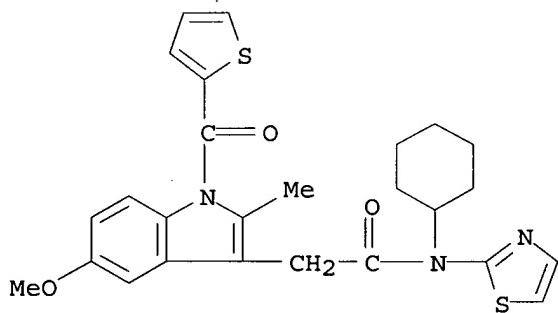
RN 773898-72-9 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-[2-(trifluoromethyl)benzoyl]- (CA INDEX NAME)



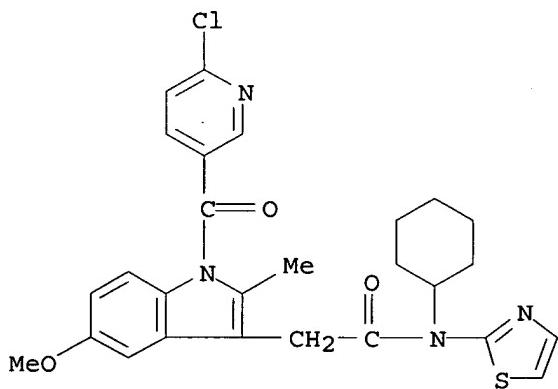
RN 773898-73-0 HCAPLUS

CN 1H-Indole-3-acetamide, N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl-1-(2-thienylcarbonyl)- (CA INDEX NAME)



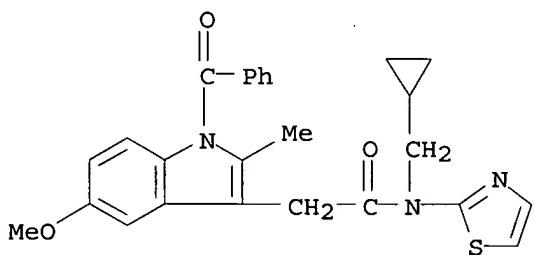
RN 773898-75-2 HCAPLUS

CN 1H-Indole-3-acetamide, 1-[(6-chloro-3-pyridinyl)carbonyl]-N-cyclohexyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



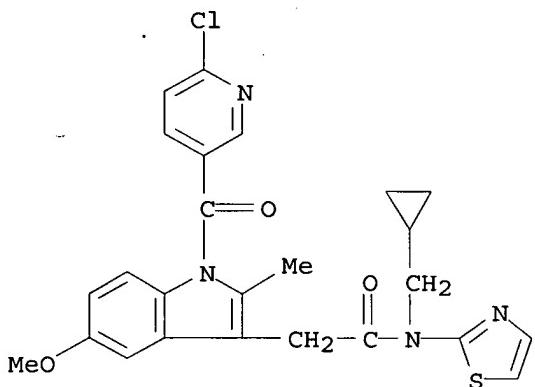
RN 773898-77-4 HCAPLUS

CN 1H-Indole-3-acetamide, 1-benzoyl-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



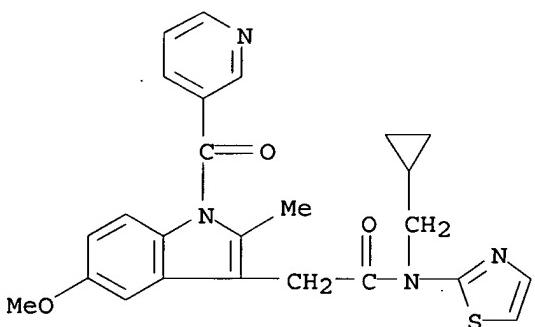
RN 773898-78-5 HCAPLUS

CN 1H-Indole-3-acetamide, 1-[(6-chloro-3-pyridinyl)carbonyl]-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



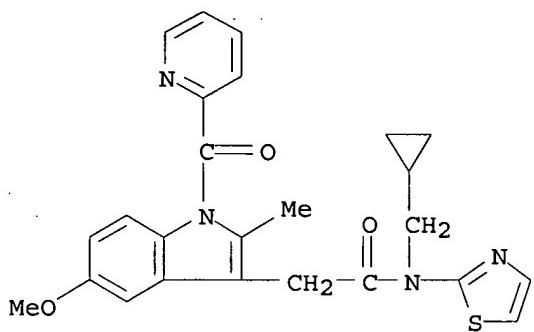
RN 773898-79-6 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(3-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



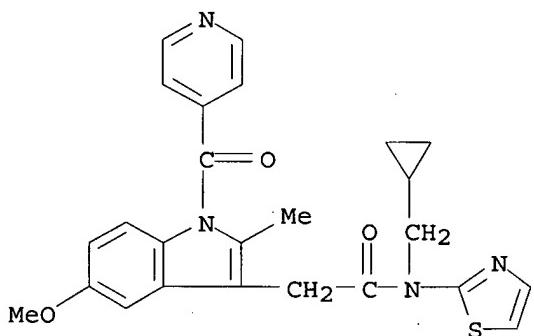
RN 773898-80-9 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(2-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



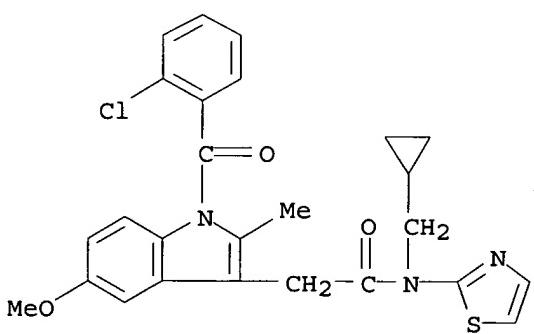
RN 773898-81-0 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(4-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



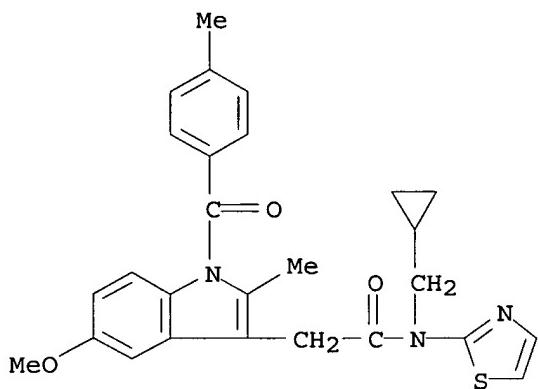
RN 773898-82-1 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(2-chlorobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



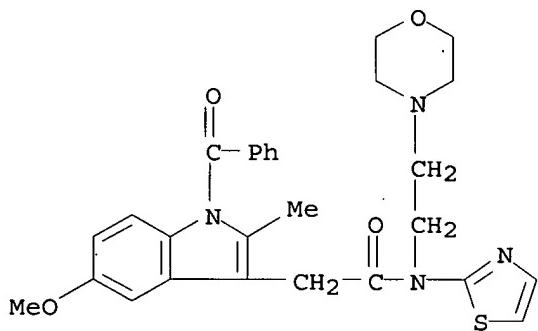
RN 773898-83-2 HCAPLUS

CN 1H-Indole-3-acetamide, N-(cyclopropylmethyl)-5-methoxy-2-methyl-1-(4-methylbenzoyl)-N-2-thiazolyl- (CA INDEX NAME)



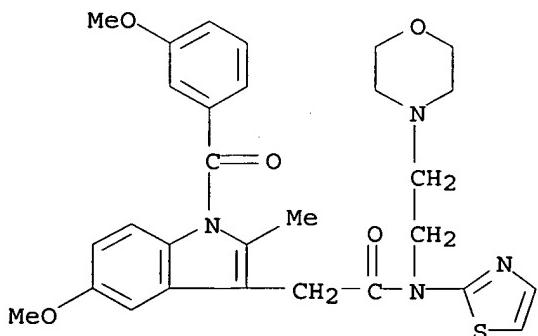
RN 773898-84-3 HCAPLUS

CN 1H-Indole-3-acetamide, 1-benzoyl-5-methoxy-2-methyl-N-[2-(4-morpholinyl)ethyl]-N-2-thiazolyl- (CA INDEX NAME)



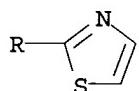
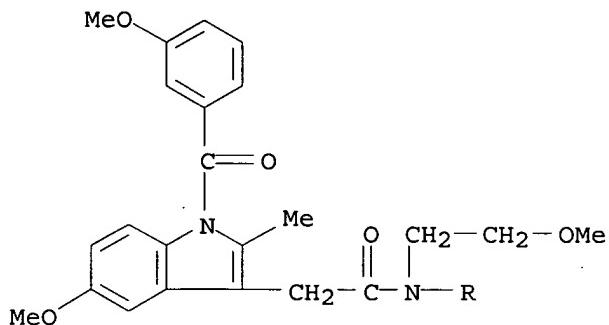
RN 773898-85-4 HCAPLUS

CN 1H-Indole-3-acetamide, 5-methoxy-1-(3-methoxybenzoyl)-2-methyl-N-[2-(4-morpholinyl)ethyl]-N-2-thiazolyl- (CA INDEX NAME)



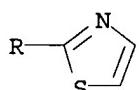
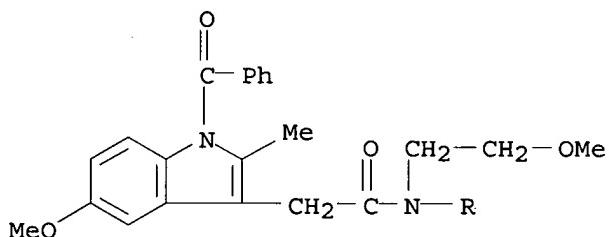
RN 773898-86-5 HCAPLUS

CN 1H-Indole-3-acetamide, 5-methoxy-1-(3-methoxybenzoyl)-N-(2-methoxyethyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



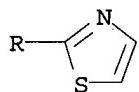
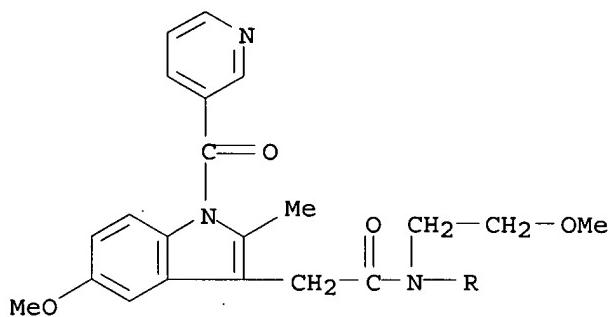
RN 773898-87-6 HCAPLUS

CN 1H-Indole-3-acetamide, 1-benzoyl-5-methoxy-N-(2-methoxyethyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



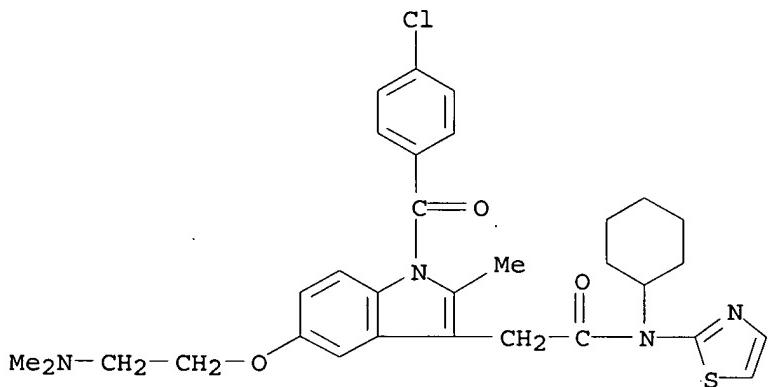
RN 773898-88-7 HCAPLUS

CN 1H-Indole-3-acetamide, 5-methoxy-N-(2-methoxyethyl)-2-methyl-1-(3-pyridinylcarbonyl)-N-2-thiazolyl- (CA INDEX NAME)



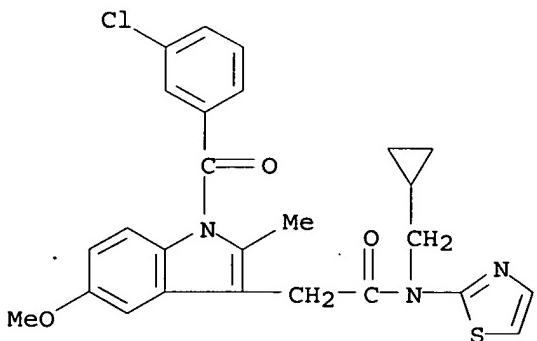
RN 773898-90-1 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-cyclohexyl-5-[2-(dimethylamino)ethoxy]-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



RN 773898-91-2 HCAPLUS

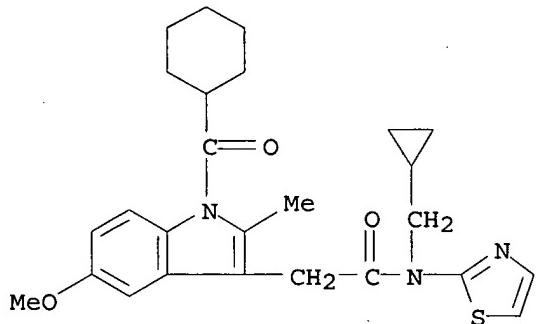
CN 1H-Indole-3-acetamide, 1-(3-chlorobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



10542169.trn

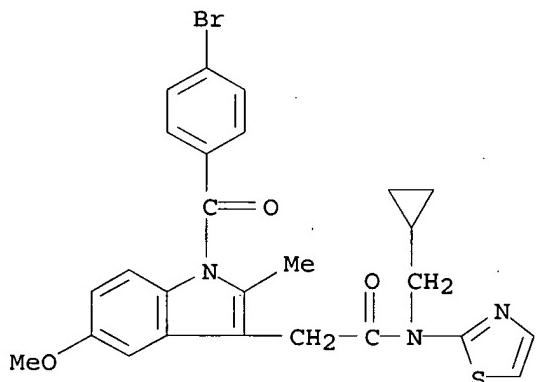
RN 773898-92-3 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(cyclohexylcarbonyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



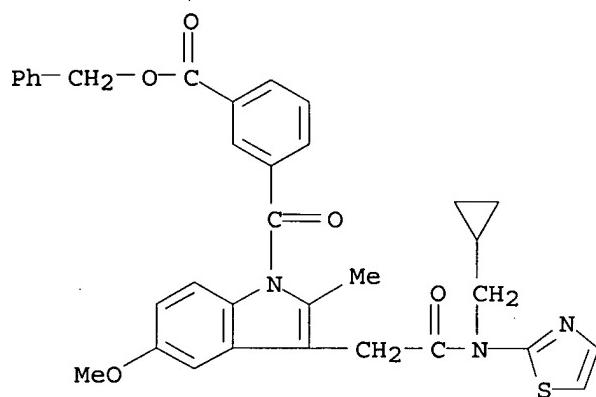
RN 773898-93-4 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-bromobenzoyl)-N-(cyclopropylmethyl)-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



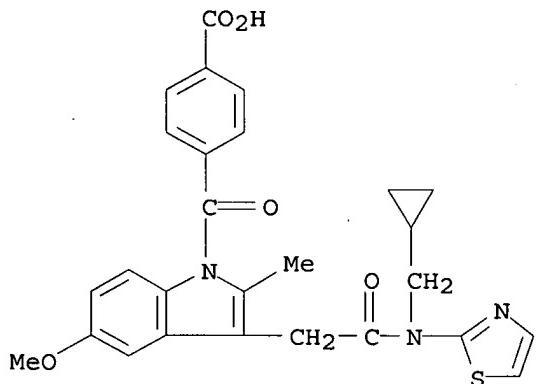
RN 773898-94-5 HCAPLUS

CN Benzoic acid, 3-[[3-[[2-[(cyclopropylmethyl)-2-thiazolylamino]-2-oxoethyl]-5-methoxy-2-methyl-1H-indol-1-yl]carbonyl]-, phenylmethyl ester (CA INDEX NAME)



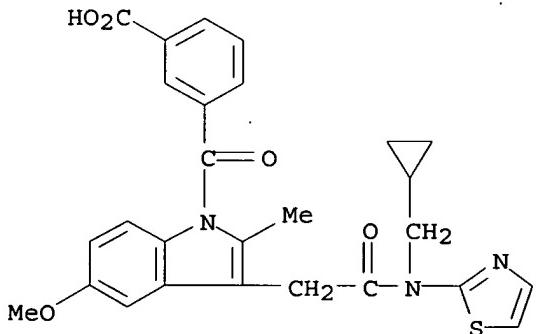
RN 773898-97-8 HCAPLUS

CN Benzoic acid, 4-[(3-[(2-[(cyclopropylmethyl)-2-thiazolylamino]-2-oxoethyl]-5-methoxy-2-methyl-1H-indol-1-yl]carbonyl]- (CA INDEX NAME)



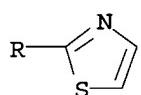
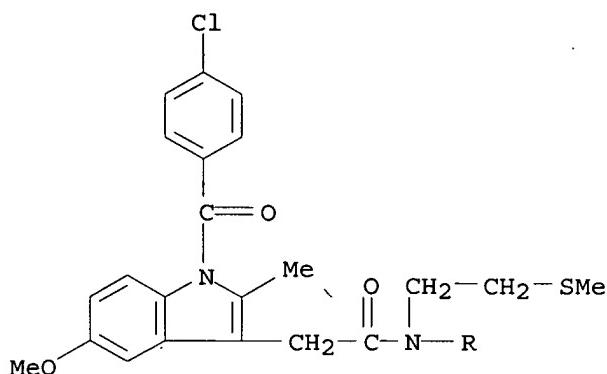
RN 773898-98-9 HCAPLUS

CN Benzoic acid, 3-[(3-[(2-[(cyclopropylmethyl)-2-thiazolylamino]-2-oxoethyl]-5-methoxy-2-methyl-1H-indol-1-yl]carbonyl]- (CA INDEX NAME)



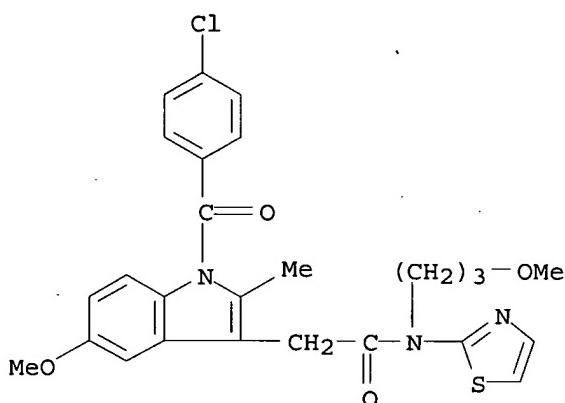
RN 773898-99-0 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[2-(methylthio)ethyl]-N-2-thiazolyl- (CA INDEX NAME)



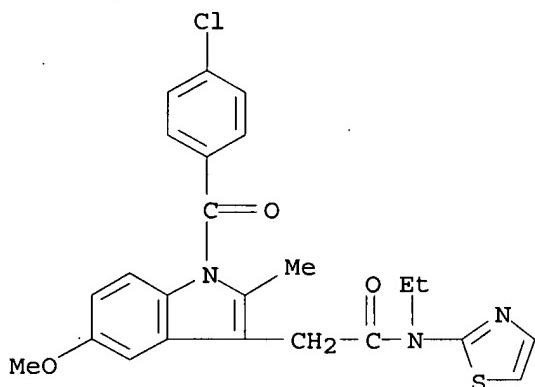
RN 773899-00-6 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-N-(3-methoxypropyl)-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



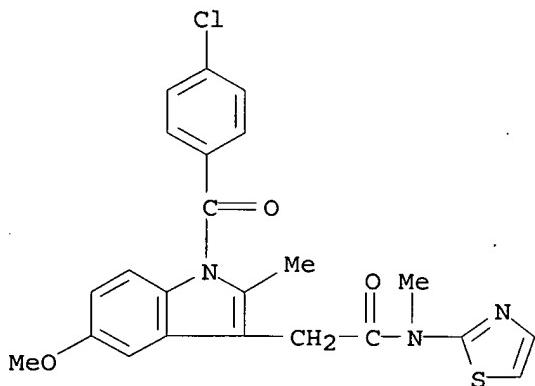
RN 773899-01-7 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-ethyl-5-methoxy-2-methyl-N-2-thiazolyl- (CA INDEX NAME)



RN 773899-02-8 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-N,2-dimethyl-2-thiazolyl- (CA INDEX NAME)



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L122 NOT FOUND

The L-number entered has not been defined in this session, or it has been deleted. To see the L-numbers currently defined in this session, enter DISPLAY HISTORY at an arrow prompt (=>).

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L22 ANSWER 1 OF 4 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2001:721438 HCPLUS

DOCUMENT NUMBER: 135:288343

TITLE: Preparation and activity of nitrosated and

nitrosylated nonsteroidal antiinflammatory compounds
Bandarage, Upul K.; Dong, Qing; Fang, Xinqin; Garvey, David S.; Mercer, Gregory J.; Richardson, Stewart K.; Schroeder, Joseph D.; Wang, Tiansheng

PATENT ASSIGNEE(S): Nitromed, Inc., USA

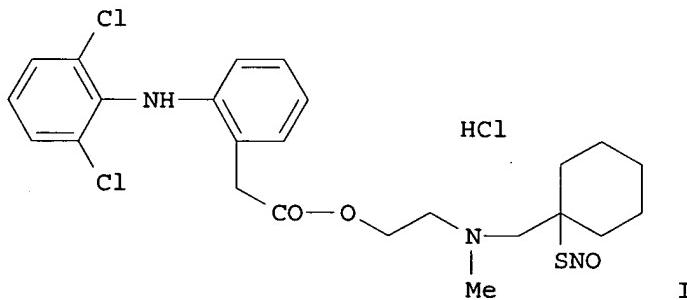
SOURCE: U.S., 59 pp., Cont.-in-part of U.S. Ser. No. 182,433, abandoned.

CODEN: USXXAM

DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|--------------|
| US 6297260 | B1 | 20011002 | US 1999-429019 | 19991029 <-- |
| CA 2348741 | A1 | 20000511 | CA 1999-2348741 | 19991029 <-- |
| WO 2000025776 | A1 | 20000511 | WO 1999-US25481 | 19991029 <-- |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AM, AZ, BY, KG, KZ,
MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE,
IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML,
MR, NE, SN, TD, TG | | | | |
| EP 1126838 | A1 | 20010829 | EP 1999-958708 | 19991029 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO | | | | |
| JP 2002528495 | T | 20020903 | JP 2000-579217 | 19991029 <-- |
| AU 763000 | B2 | 20030710 | AU 2000-16012 | 19991029 <-- |
| US 2002016322 | A1 | 20020207 | US 2001-938560 | 20010827 <-- |
| US 6593347 | B2 | 20030715 | | |
| US 2003207919 | A1 | 20031106 | US 2003-431457 | 20030508 <-- |
| AU 2004200091 | A1 | 20040205 | AU 2004-200091 | 20040109 |
| PRIORITY APPLN. INFO.: | | | US 1998-182433 | B2 19981030 |
| | | | AU 2000-16012 | A 19991029 |
| | | | US 1999-429019 | A3 19991029 |
| | | | WO 1999-US25481 | W 19991029 |
| | | | US 2001-938560 | A3 20010827 |

OTHER SOURCE(S) : MARPAT 135:288343
 GI



AB The present invention describes novel nitrosated and/or nitrosylated nonsteroidal antiinflammatory compds., and novel compns. comprising at least one nitrosated and/or nitrosylated nonsteroidal antiinflammatory compound, and, optionally, at least one compound that donates, transfers or releases nitric oxide, elevates endogenous levels of endothelium-derived relaxing factor, stimulates endogenous synthesis of nitric oxide or is a substrate for nitric oxide synthase. The present invention also provides methods for treating, preventing and/or reducing inflammation, pain, and fever; decreasing or reversing the gastrointestinal, renal and

other toxicities resulting from the use of nonsteroidal antiinflammatory drugs; treating and/or preventing gastrointestinal disorders; treating inflammatory disease states and disorders; and treating and/or preventing ophthalmic diseases or disorders.

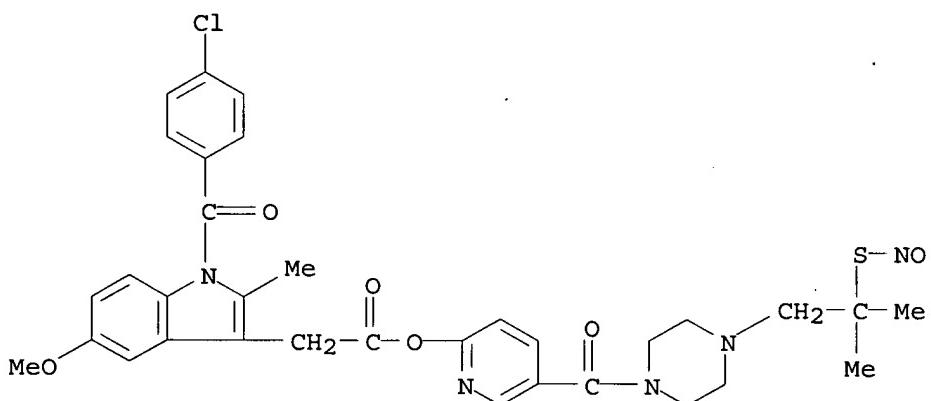
Thus, I was prepared in 8 steps from cyclohexanecarboxaldehyde and shows a relative activity of 1, 1.2 and 0.02 in analgesic, antiinflammatory and gastric lesion tests.

IT 364590-30-7P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation and activity of nitrosated and nitrosylated nonsteroidal antiinflammatory compds.)

RN 364590-30-7 HCPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-, 5-[[4-[2-methyl-2-(nitrosothio)propyl]-1-piperazinyl]carbonyl]-2-pyridinyl ester (CA INDEX NAME)



REFERENCE COUNT: 63 THERE ARE 63 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 2 OF 4 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2000:314687 HCPLUS

DOCUMENT NUMBER: 132:334454

TITLE: Preparation of 2-amino-thiazole derivatives as antitumor agents

INVENTOR(S): Pevarello, Paolo; Amici, Raffaella; Traquandi, Gabriella; Villa, Manuela; Vulpetti, Anna; Isacchi, Antonella

PATENT ASSIGNEE(S): Pharmacia & Upjohn S.p.A., Italy

SOURCE: PCT Int. Appl., 115 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

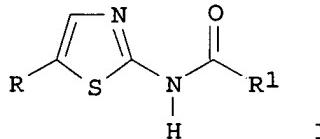
| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|--------------|
| WO 2000026202 | A1 | 20000511 | WO 1999-EP8306 | 19991027 <-- |
| W: AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM, | | | | |

| | | | | |
|---|----|----------|------------------|--------------|
| AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, | | | | |
| DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, | | | | |
| CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2347188 | A1 | 20000511 | CA 1999-2347188 | 19991027 <-- |
| AU 200012679 | A | 20000522 | AU 2000-12679 | 19991027 <-- |
| AU 766193 | B2 | 20031009 | | |
| EP 1124810 | A1 | 20010822 | EP 1999-955931 | 19991027 <-- |
| EP 1124810 | B1 | 20050504 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, | | | | |
| IE, SI, LT, LV, FI, RO | | | | |
| BR 9914958 | A | 20011218 | BR 1999-14958 | 19991027 <-- |
| HU 2001004200 | A2 | 20020328 | HU 2001-4200 | 19991027 <-- |
| HU 2001004200 | A3 | 20031229 | | |
| JP 2002528537 | T | 20020903 | JP 2000-579591 | 19991027 <-- |
| NZ 510965 | A | 20031031 | NZ 1999-510965 | 19991027 <-- |
| TW 222447 | B | 20041021 | TW 1999-88118558 | 19991027 |
| AT 294785 | T | 20050515 | AT 1999-955931 | 19991027 |
| PT 1124810 | T | 20050930 | PT 1999-955931 | 19991027 |
| ES 2241338 | T3 | 20051016 | ES 1999-955931 | 19991027 |
| ZA 2001002870 | A | 20011010 | ZA 2001-2870 | 20010406 <-- |
| NO 2001002057 | A | 20010628 | NO 2001-2057 | 20010426 <-- |
| US 7037929 | B1 | 20060502 | US 2001-807962 | 20010426 <-- |
| MX 2001PA04278 | A | 20020621 | MX 2001-PA4278 | 20010427 <-- |
| IN 2001CN00744 | A | 20050304 | IN 2001-CN744 | 20010528 |
| AU 2004200096 | A1 | 20040205 | AU 2004-200096 | 20040109 |
| PRIORITY APPLN. INFO.: | | | GB 1998-23871 | A 19981030 |
| | | | US 1998-823871 | A 19981030 |
| | | | AU 2000-12679 | A 19991027 |
| | | | WO 1999-EP8306 | W 19991027 |

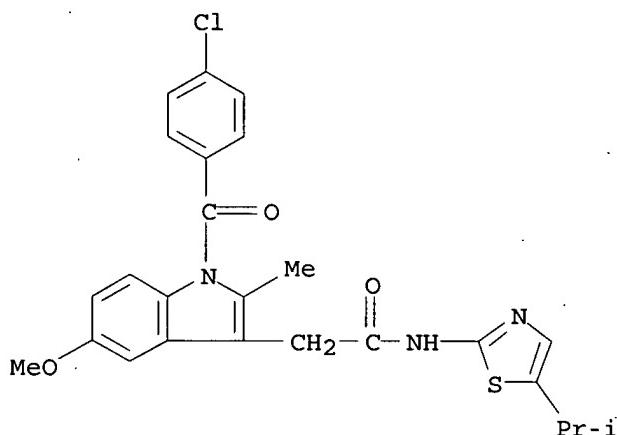
OTHER SOURCE(S) :

MARPAT 132:334454

GI



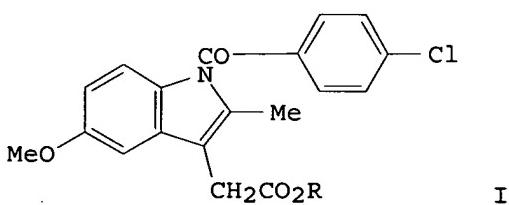
- AB The title compds. [I; R = halo, NO₂, (un)substituted amino NH₂, etc.; R₁ = alkyl, alkenyl, 3-6 membered carbocycle, etc.], useful for treating cell proliferative disorders associated with an altered cell dependent kinase activity such as cancer, Alzheimer's disease, viral infections, autoimmune diseases or neurodegenerative disorders, were prepared E.g., thiazole I [R = iso-Pr; R₁ = 4-Me₂NC₆H₄CH₂] showed Ki of 0.1 μM against cdk2/cyclin A complex.
- IT 267656-89-3P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of 2-amino-thiazole derivs. as antitumor agents)
- RN 267656-89-3 HCPLUS
- CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[5-(1-methylethyl)-2-thiazolyl]- (CA INDEX NAME)



REFERENCE COUNT: 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 3 OF 4 HCAPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 1980:639224 HCAPLUS
 DOCUMENT NUMBER: 93:239224
 ORIGINAL REFERENCE NO.: 93:38315a, 38318a
 TITLE: Pharmaceutical 1-(p-chlorobenzoyl)-5-methoxy-2-methylindole-3-acet-3-oxy-1-isobenzofuranyl esters
 INVENTOR(S): Vandoni, Guido
 PATENT ASSIGNEE(S): Resfar S.r.l., Italy
 SOURCE: Ger. Offen., 18 pp.
 CODEN: GWXXBX
 DOCUMENT TYPE: Patent
 LANGUAGE: German
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|--------------|
| DE 3005827 | A1 | 19800904 | DE 1980-3005827 | 19800216 <-- |
| US 4277489 | A | 19810707 | US 1980-119332 | 19800207 <-- |
| JP 55113778 | A | 19800902 | JP 1980-19817 | 19800221 <-- |
| JP 59051954 | B | 19841217 | | |
| FR 2449686 | A1 | 19800919 | FR 1980-3865 | 19800221 <-- |
| FR 2449686 | B1 | 19860425 | | |
| PRIORITY APPLN. INFO.: | | | IT 1979-20398 | A 19790221 |
| GI | | | | |



I

AB Indole I ($R = \text{phthalidyl}$) (II) was prepared by treating I ($R = \text{H}$) with bromophthalide in $\text{Et}_3\text{N}/\text{CHCl}_3$. Tests showed that II had higher

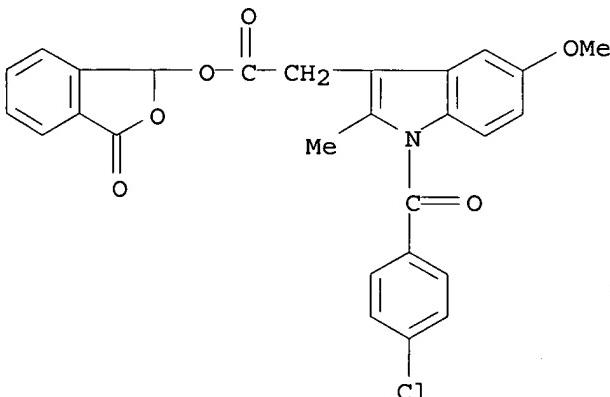
antiphlogistic and analgesic activities than I ($R = H$), with lower ulcerogenic activity.

IT 67489-39-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation and analgesic and antiinflammatory activity of)

RN 67489-39-8 HCAPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-, 1,3-dihydro-3-oxo-1-isobenzofuranyl ester (CA INDEX NAME)



L22 ANSWER 4 OF 4 HCAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1970:78871 HCAPLUS

DOCUMENT NUMBER: 72:78871

ORIGINAL REFERENCE NO.: 72:14361a,14364a

TITLE: 1-(p-Chlorobenzoyl)-2-formyl-3-indolyl acetic acids

INVENTOR(S): Chemerda, John M.; Sletzinger, Meyer

PATENT ASSIGNEE(S): Merck and Co., Inc.

SOURCE: U.S., 3 pp.

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|--------------|
| US 3489765 | A | 19700113 | US 1967-656008 | 19670726 <-- |
| PRIORITY APPLN. INFO.: | | | US 1967-656008 | A 19670726 |

GI For diagram(s), see printed CA Issue.

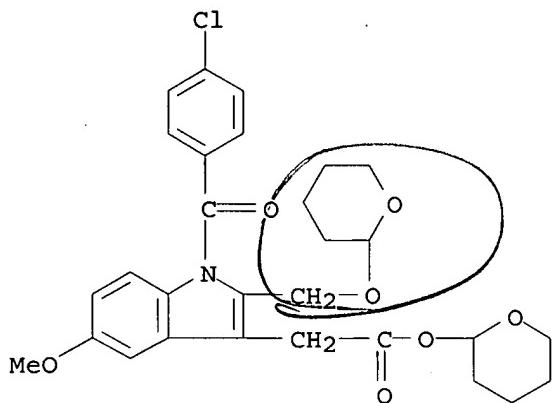
AB Compds. (I) were prepared by treating II with N_2H_4 or $NH_2CONHNH_2$ and reducing the hydrazone or semicarbazone. Thus 3.72 g II ($R = MeO$) and 0.64 g NH_2NH_2 gave 4.19 g hydrazone, which was reduced with Me_3COK - Me_2SO to give I ($R = OMe$). Similarly prepared was I ($R = Me_2N$).

IT 25998-02-1P 26129-01-1P

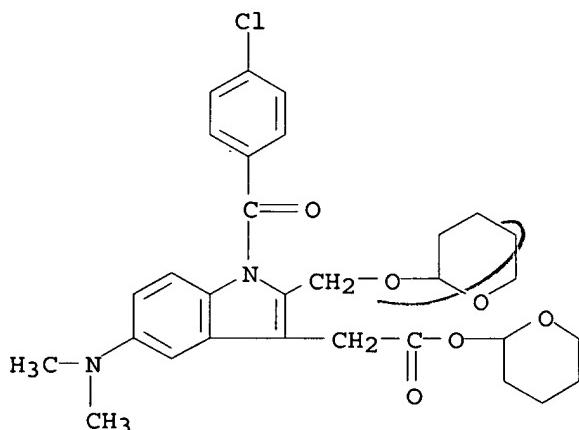
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 25998-02-1 HCAPLUS

CN Indole-3-acetic acid, 1-(p-chlorobenzoyl)-5-methoxy-2-[(tetrahydro-2H-pyran-2-yl)oxy]methyl-, tetrahydro-2H-pyran-2-yl ester (8CI) (CA INDEX NAME)



RN 26129-01-1 HCAPLUS

CN Indole-3-acetic acid, 1-(p-chlorobenzoyl)-5-(dimethylamino)-2-[(tetrahydro-2H-pyran-2-yl)oxymethyl]-, tetrahydro-2H-pyran-2-yl ester
(8CI) (CA INDEX NAME)

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L23 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2003:377132 HCAPLUS

DOCUMENT NUMBER: 138:367144

TITLE: Soluble CD40L (CD154) as a prognostic marker of atherosclerotic diseases

INVENTOR(S): Schoenbeck, Uwe; Ridker, Paul M.; Libby, Peter

PATENT ASSIGNEE(S): The Brigham and Women's Hospital, Inc., USA

SOURCE: PCT Int. Appl., 66 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------|-------|-------|-----------------|-------|
| ----- | ----- | ----- | ----- | ----- |

| | | | | |
|--|----|----------|-----------------|--------------|
| WO 2003040691 | A2 | 20030515 | WO 2002-US35505 | 20021105 <-- |
| WO 2003040691 | A3 | 20031113 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
UA, UG, UZ, VN, YU, ZA, ZM, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF,
CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2464531 | A1 | 20030515 | CA 2002-2464531 | 20021105 <-- |
| AU 2002343620 | A1 | 20030519 | AU 2002-343620 | 20021105 <-- |
| US 2003152566 | A1 | 20030814 | US 2002-288253 | 20021105 <-- |
| US 7189518 | B2 | 20070313 | | |
| EP 1451577 | A2 | 20040901 | EP 2002-780578 | 20021105 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK | | | | |
| CN 1613012 | A | 20050504 | CN 2002-826711 | 20021105 |
| JP 2005515407 | T | 20050526 | JP 2003-542897 | 20021105 |
| PRIORITY APPLN. INFO.: | | | US 2001-338841P | P 20011105 |
| | | | WO 2002-US35505 | W 20021105 |

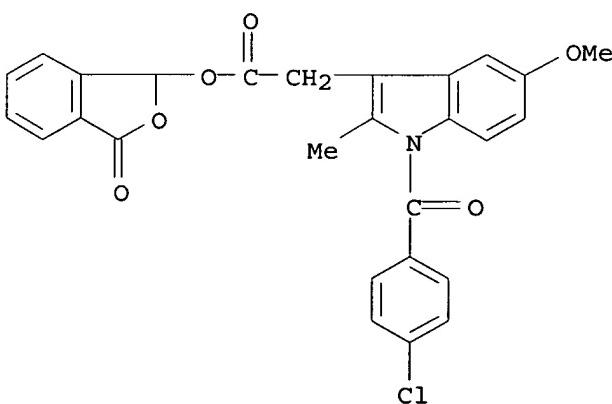
AB The invention involves the new use of a diagnostic test to determine the risk of atherosclerotic diseases, e.g. myocardial infarction and stroke, particularly among individuals with no signs or symptoms of current disease and among nonsmokers. Further, the invention involves the new use of a diagnostic test to assist physicians in determining which individuals at risk will preferentially benefit from certain treatments designed either to prevent first or recurrent myocardial infarctions and strokes, or to treat acute and chronic cardiovascular disorders. Methods for treatment are also described.

IT 67489-39-8, Talmetacin

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(soluble CD40L as prognostic marker of atherosclerotic diseases, and use in therapeutic agent assessment)

RN 67489-39-8 HCAPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-,
1,3-dihydro-3-oxo-1-isobenzofuranyl ester (CA INDEX NAME)

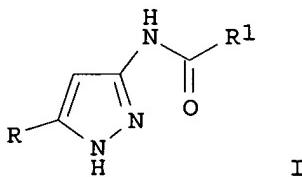


=> d 124 ibib abs hitstr tot

L24 ANSWER 1 OF 10 HCAPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 2006:374223 HCAPLUS
 DOCUMENT NUMBER: 144:412501
 TITLE: Preparation of 3(5)-acylaminopyrazole derivatives for use as therapeutic agents, particularly antitumor agents
 INVENTOR(S): Pevarello, Paolo; Orsini, Paolo; Traquandi, Gabriella; Varasi, Mario; Fritzen, Edward L.; Warpehoski, Martha A.; Pierce, Betsy S.; Brasca, Maria Grabriella
 PATENT ASSIGNEE(S): Pharmacia Italia S.p.A., Italy; Pharmacia & Upjohn Company LLC
 SOURCE: U.S., 41 pp., Cont.-in-part of U.S. Ser. No. 372,831, abandoned.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| <u>US 7034049</u> | B1 | 20060425 | US 2002-48486 | 20020501 <-- |
| <u>WO 2001012189</u> | A1 | 20010222 | WO 2000-US6699 | 20000505 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| <u>US 6218418</u> | B1 | 20010417 | US 2000-667603 | 20000922 <-- |
| PRIORITY APPLN. INFO.: | | | US 1999-372831 | B2 19990812 |
| | | | WO 2000-US6699 | W 20000505 |
| | | | US 2000-560400 | A1 20000428 |

OTHER SOURCE(S): MARPAT 144:412501
 GI



AB Compds. (e.g., N-(5-cyclopropyl-1H-pyrazol-3-yl)-2,2-diphenylacetamide) which are 3-amino-pyrazole derivs. represented by formula I (wherein R = C3-C6 cycloalkyl group optionally substituted by a straight or branched C1-C6 alkyl or arylalkyl group; R1 = a straight or branched C1-C6 alkyl, C2-C4 alkenyl, cycloalkyl, cycloalkenyl, heterocyclyl, aryl, arylalkyl, arylcarbonyl, aryloxyalkyl or arylalkenyl group, each of which may be optionally further substituted) are claimed. A process for preparing the 3-aminopyrazole derivs. comprises: (a) reacting RCO2R2 (R2 = alkyl), with

MeCN in the presence of a basic agent, to obtain $\text{RC(O)CH}_2\text{CN}$; (b) reacting $\text{RC(O)CH}_2\text{CN}$ with hydrazine hydrate to obtain an 3-amino-5-R-1H-pyrazole; (c) oxidizing the 3-amino-5-R-1H-pyrazole to obtain the nitro analog; (d) reacting the nitro compound with tert-butoxycarbonyl anhydride (Boc_2O) to obtain the N-Boc derivative which was reduced; (e) reacting this amino compound with $\text{R}_1\text{C(O)X}$ ($\text{X} = \text{OH}$ or a suitable leaving group) to obtain the N1-Boc-protected I; and (g) hydrolyzing this intermediate in an acidic medium to obtain I. The compds. are useful for the treatment of cancer, cell proliferative disorders, Alzheimer's disease, viral infections, auto-immune diseases or neurodegenerative diseases (no data is given). Pharmaceutical compns. containing I are also claimed.

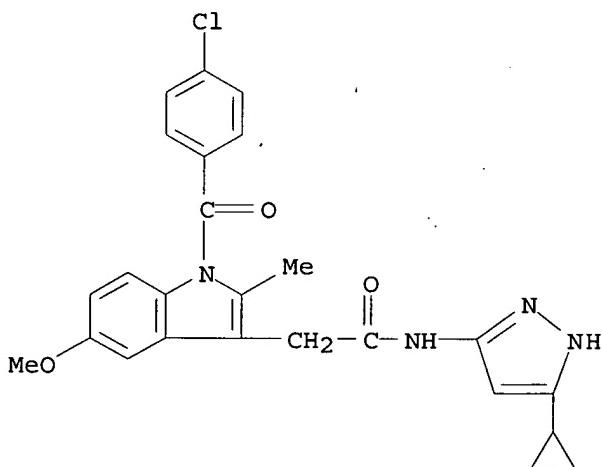
IT 326824-45-7P, 2-[1-(4-Chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-cyclopropyl-1H-pyrazol-3-yl)acetamide

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of 3(5)-acylaminopyrazole derivs. for use as therapeutic agents, particularly antitumor agents)

RN 326824-45-7 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-(5-cyclopropyl-1H-pyrazol-3-yl)-5-methoxy-2-methyl- (CA INDEX NAME)



REFERENCE COUNT: 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 2 OF 10 HCPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2003:377132 HCPLUS

DOCUMENT NUMBER: 138:367144

TITLE: Soluble CD40L (CD154) as a prognostic marker of atherosclerotic diseases

INVENTOR(S): Schoenbeck, Uwe; Ridker, Paul M.; Libby, Peter

PATENT ASSIGNEE(S): The Brigham and Women's Hospital, Inc., USA

SOURCE: PCT Int. Appl., 66 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.

KIND

DATE

APPLICATION NO.

DATE

| | | | | |
|------------------------|--|----------|-----------------|--------------|
| ----- | ----- | ----- | ----- | |
| WO 2003040691 | A2 | 20030515 | WO 2002-US35505 | 20021105 <-- |
| WO 2003040691 | A3 | 20031113 | | |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | |
| CA 2464531 | A1 | 20030515 | CA 2002-2464531 | 20021105 <-- |
| AU 2002343620 | A1 | 20030519 | AU 2002-343620 | 20021105 <-- |
| US 2003152566 | A1 | 20030814 | US 2002-288253 | 20021105 <-- |
| US 7189518 | B2 | 20070313 | | |
| EP 1451577 | A2 | 20040901 | EP 2002-780578 | 20021105 |
| R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK | | | |
| CN 1613012 | A | 20050504 | CN 2002-826711 | 20021105 |
| JP 2005515407 | T | 20050526 | JP 2003-542897 | 20021105 |
| PRIORITY APPLN. INFO.: | | | US 2001-338841P | P 20011105 |
| | | | WO 2002-US35505 | W 20021105 |

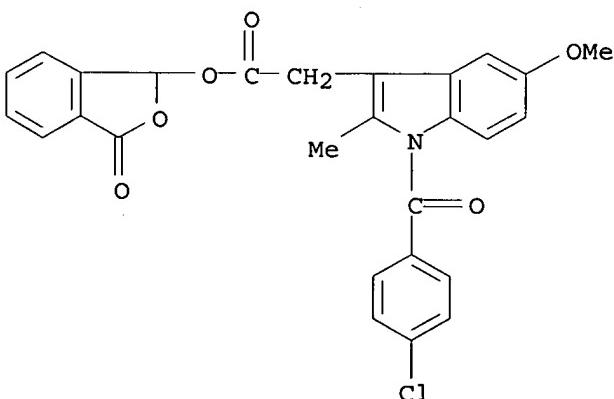
AB The invention involves the new use of a diagnostic test to determine the risk of atherosclerotic diseases, e.g. myocardial infarction and stroke, particularly among individuals with no signs or symptoms of current disease and among nonsmokers. Further, the invention involves the new use of a diagnostic test to assist physicians in determining which individuals at risk will preferentially benefit from certain treatments designed either to prevent first or recurrent myocardial infarctions and strokes, or to treat acute and chronic cardiovascular disorders. Methods for treatment are also described.

IT 67489-39-8, Talmetacin

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(soluble CD40L as prognostic marker of atherosclerotic diseases, and use in therapeutic agent assessment)

RN 67489-39-8 HCPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-, 1,3-dihydro-3-oxo-1-isobenzofuranyl ester (CA INDEX NAME)



L24 ANSWER 3 OF 10 HCAPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 2001:780679 HCAPLUS
 DOCUMENT NUMBER: 135:327362
 TITLE: Nonsteroidal antiinflammatory drug (NSAID) and NSAID derivative amyloid A β 42 polypeptide-lowering agents for the treatment of Alzheimer's disease, and screening methods
 INVENTOR(S): Koo, Edward Hao Mang; Golde, Todd Eliot; Galasko, Douglas Roger
 PATENT ASSIGNEE(S): Mayo Foundation for Medical Education and Research, USA
 SOURCE: PCT Int. Appl., 73 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|--------------|
| WO 2001078721 | A1 | 20011025 | WO 2001-US11956 | 20010412 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2406383 | A1 | 20011025 | CA 2001-2406383 | 20010412 <-- |
| AU 200157022 | A | 20011030 | AU 2001-57022 | 20010412 <-- |
| EP 1284729 | A1 | 20030226 | EP 2001-930491 | 20010412 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | | |
| JP 2003530437 | T | 20031014 | JP 2001-576021 | 20010412 <-- |
| US 2002128319 | A1 | 20020912 | US 2001-12606 | 20011207 <-- |
| US 6911466 | B2 | 20050628 | | |
| US 2005089945 | A1 | 20050428 | US 2004-928925 | 20040827 <-- |
| US 7097998 | B2 | 20060829 | | |
| US 2005186559 | A1 | 20050825 | US 2005-113789 | 20050425 <-- |
| AU 2005201819 | A1 | 20050519 | AU 2005-201819 | 20050429 |
| AU 2005201819 | B2 | 20070712 | | |
| US 2006004086 | A1 | 20060105 | US 2005-170776 | 20050628 <-- |
| US 2007253905 | A1 | 20071101 | US 2007-740791 | 20070426 <-- |
| US 2007253906 | A1 | 20071101 | US 2007-740800 | 20070426 <-- |
| AU 2007224395 | A1 | 20071101 | AU 2007-224395 | 20071011 |
| PRIORITY APPLN. INFO.: | | | US 2000-196617P | P 20000413 |
| | | | AU 2001-257022 | A3 20010412 |
| | | | WO 2001-US11956 | W 20010412 |
| | | | US 2001-12606 | A3 20011207 |
| | | | US 2005-113789 | A1 20050425 |
| | | | AU 2005-201819 | A3 20050429 |

AB A method is provided for preventing, delaying, or reversing the progression of Alzheimer's disease by administering an A β 42-lowering agent to a mammal under conditions in which levels of A β 42 are selectively reduced, levels of A β 38 are increased, and levels of A β 40 are unchanged. The invention provides methods and materials for developing and identifying A β 42-lowering agents. In addition, the invention provides methods for identifying agents that increase the risk

of developing, or hasten progression of, Alzheimer's disease. The invention also provides compns. of A β 42-lowering agents and antioxidants, A β 42 lowering agents and non-selective secretase inhibitors, and A β 42 lowering agents and acetylcholinesterase inhibitors. The invention further provides kits containing A β 42-lowering agents, antioxidants, non-selective secretase inhibitors, and/or acetylcholinesterase inhibitors as well as instructions related to dose regimens for A β 42-lowering agents, antioxidants, non-selective secretase inhibitors, and acetylcholinesterase inhibitors. The agents of the invention include nonsteroidal antiinflammatory drugs (NSAIDs) and NSAID derivs.

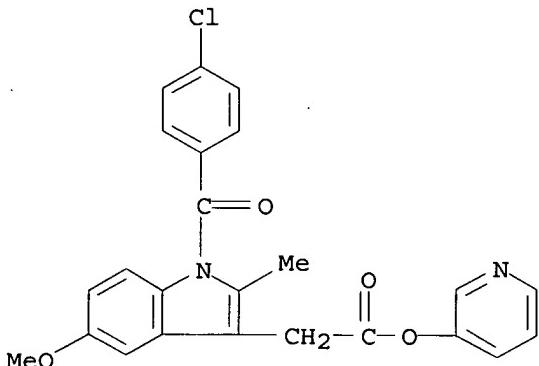
IT 80590-83-6 261766-29-4

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(NSAID and NSAID derivative amyloid A β 42 polypeptide-lowering agents for treatment of Alzheimer's disease, and screening methods)

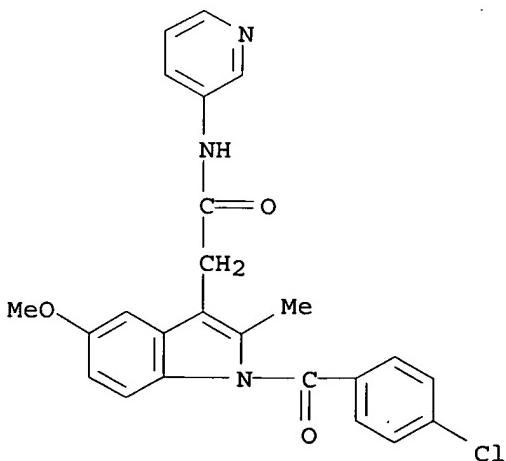
RN 80590-83-6 HCPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-, 3-pyridinyl ester (CA INDEX NAME)



RN 261766-29-4 HCPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-3-pyridinyl- (CA INDEX NAME)

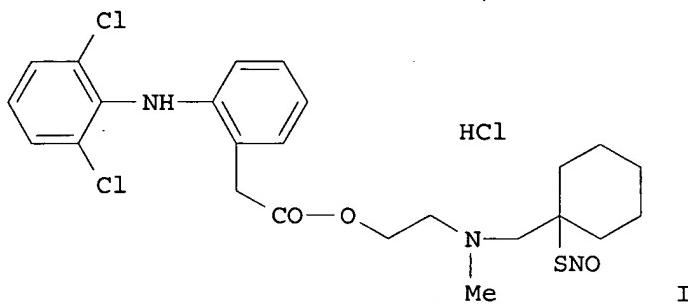


REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 4 OF 10 HCAPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 2001:721438 HCAPLUS
 DOCUMENT NUMBER: 135:288343
 TITLE: Preparation and activity of nitrosated and
 nitrosylated nonsteroidal antiinflammatory compounds
 INVENTOR(S): Bandarage, Upul K.; Dong, Qing; Fang, Xinqin; Garvey,
 David S.; Mercer, Gregory J.; Richardson, Stewart K.;
 Schroeder, Joseph D.; Wang, Tiansheng
 PATENT ASSIGNEE(S): Nitromed, Inc., USA
 SOURCE: U.S., 59 pp., Cont.-in-part of U.S. Ser. No. 182,433,
 abandoned.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| US 6297260 | B1 | 20011002 | US 1999-429019 | 19991029 <-- |
| CA 2348741 | A1 | 20000511 | CA 1999-2348741 | 19991029 <-- |
| WO 2000025776 | A1 | 20000511 | WO 1999-US25481 | 19991029 <-- |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AM, AZ, BY, KG, KZ,
MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE,
IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML,
MR, NE, SN, TD, TG | | | | |
| EP 1126838 | A1 | 20010829 | EP 1999-958708 | 19991029 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO | | | | |
| JP 2002528495 | T | 20020903 | JP 2000-579217 | 19991029 <-- |
| AU 763000 | B2 | 20030710 | AU 2000-16012 | 19991029 <-- |
| US 2002016322 | A1 | 20020207 | US 2001-938560 | 20010827 <-- |
| US 6593347 | B2 | 20030715 | | |
| US 2003207919 | A1 | 20031106 | US 2003-431457 | 20030508 <-- |
| AU 2004200091 | A1 | 20040205 | AU 2004-200091 | 20040109 |
| PRIORITY APPLN. INFO.: | | | US 1998-182433 | B2 19981030 |
| | | | AU 2000-16012 | A 19991029 |
| | | | US 1999-429019 | A3 19991029 |
| | | | WO 1999-US25481 | W 19991029 |
| | | | US 2001-938560 | A3 20010827 |

OTHER SOURCE(S): MARPAT 135:288343
 GI



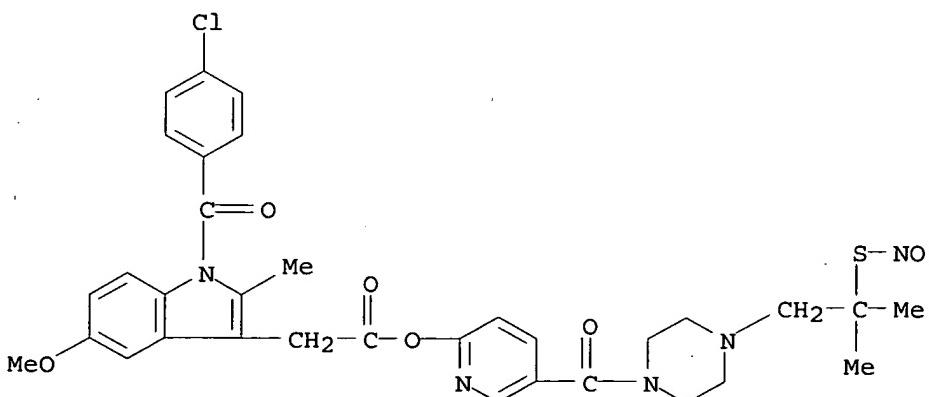
AB The present invention describes novel nitrosated and/or nitrosylated nonsteroidal antiinflammatory compds., and novel compns. comprising at least one nitrosated and/or nitrosylated nonsteroidal antiinflammatory compound, and, optionally, at least one compound that donates, transfers or releases nitric oxide, elevates endogenous levels of endothelium-derived relaxing factor, stimulates endogenous synthesis of nitric oxide or is a substrate for nitric oxide synthase. The present invention also provides methods for treating, preventing and/or reducing inflammation, pain, and fever; decreasing or reversing the gastrointestinal, renal and other toxicities resulting from the use of nonsteroidal antiinflammatory drugs; treating and/or preventing gastrointestinal disorders; treating inflammatory disease states and disorders; and treating and/or preventing ophthalmic diseases or disorders. Thus, I was prepared in 8 steps from cyclohexanecarboxaldehyde and shows a relative activity of 1, 1.2 and 0.02 in analgesic, antiinflammatory and gastric lesion tests.

IT 364590-30-7P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation and activity of nitrosated and nitrosylated nonsteroidal antiinflammatory compds.)

RN 364590-30-7 HCPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-, 5-[[4-[2-methyl-2-(nitrosothio)propyl]-1-piperazinyl]carbonyl]-2-pyridinyl ester (CA INDEX NAME)



REFERENCE COUNT:

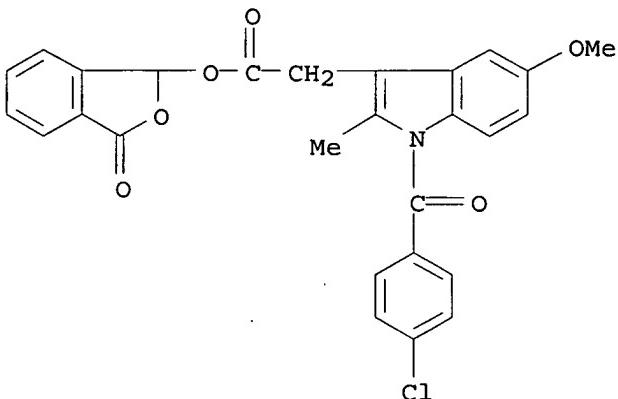
63

THERE ARE 63 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 5 OF 10 HCAPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 2001:167849 HCAPLUS
 DOCUMENT NUMBER: 134:217194
 TITLE: Systemic inflammatory markers as diagnostic tools in the prevention of atherosclerotic diseases
 INVENTOR(S): Ridker, Paul; Hennekens, Charles H.
 PATENT ASSIGNEE(S): The Brigham and Women's Hospital, Inc., USA
 SOURCE: PCT Int. Appl., 53 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 2001015744 | A1 | 20010308 | WO 2000-US24251 | 20000831 <-- |
| WO 2001015744 | A9 | 20020926 | | |
| W: AU, CA, JP | | | | |
| RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE | | | | |
| US 7030152 | B1 | 20060418 | US 1999-387028 | 19990831 <-- |
| CA 2381926 | A1 | 20010308 | CA 2000-2381926 | 20000831 <-- |
| AU 200071103 | A | 20010326 | AU 2000-71103 | 20000831 <-- |
| AU 782386 | B2 | 20050721 | | |
| EP 1212101 | A1 | 20020612 | EP 2000-959851 | 20000831 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY | | | | |
| JP 2003508453 | T | 20030304 | JP 2001-520155 | 20000831 <-- |
| AU 2005225101 | A1 | 20051117 | AU 2005-225101 | 20051021 |
| PRIORITY APPLN. INFO.: | | | US 1999-387028 | A 19990831 |
| | | | US 1997-41950P | P 19970402 |
| | | | US 1997-43039P | P 19970402 |
| | | | US 1998-70894P | P 19980109 |
| | | | US 1998-54212 | A2 19980402 |
| | | | WO 2000-US24251 | W 20000831 |

- AB The invention involves methods for characterizing an individual's risk profile of developing a future cardiovascular disorder such as atherosclerosis, stroke, and myocardial infarction by assessing the level of systemic inflammation marker (such as sICAM or C-reactive protein) in an individual. The invention also involves methods for evaluating the likelihood that an individual will benefit from treatment with an agent for reducing the risk of future cardiovascular disorders; and of drug combinations (anti-inflammatory agents, lipid-reducing agents, angiotensin system inhibitors, calcium channel blockers, β -adrenergic receptor blockers) suitable for prevention future cardiovascular disease.
- IT 67489-39-8, Talmetacin
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (use of agents and systemic inflammatory markers to predict and inhibit cardiovascular disorders in humans)
- RN 67489-39-8 HCAPLUS
- CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-, 1,3-dihydro-3-oxo-1-isobenzofuranyl ester (CA INDEX NAME)



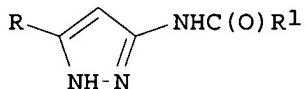
REFERENCE COUNT: 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 6 OF 10 HCAPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 2001:137023 HCAPLUS
 DOCUMENT NUMBER: 134:178552
 TITLE: 3(5)-Acylaminopyrazole derivatives, process for their preparation and their use as antitumor agents
 INVENTOR(S): Pevarello, Paolo; Orsini, Paolo; Traquandi, Gabriella; Varasi, Mario; Fritzen, Edward L.; Warpehoski, Martha A.; Pierce, Betsy S.; Brasca, Maria Grabriella
 PATENT ASSIGNEE(S): Pharmacia & Upjohn S.p.A., Italy; Pharmacia & Upjohn Company
 SOURCE: PCT Int. Appl., 123 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 2001012189 | A1 | 20010222 | WO 2000-US6699 | 20000505 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2383555 | A1 | 20010222 | CA 2000-2383555 | 20000505 <-- |
| AU 200049714 | A | 20010313 | AU 2000-49714 | 20000505 <-- |
| EP 1202733 | A1 | 20020508 | EP 2000-931906 | 20000505 <-- |
| EP 1202733 | B1 | 20051005 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL | | | | |
| BR 2000013143 | A | 20020611 | BR 2000-13143 | 20000505 <-- |
| JP 2003507329 | T | 20030225 | JP 2001-516535 | 20000505 <-- |
| EE 200200065 | A | 20030415 | EE 2002-65 | 20000505 <-- |
| HU 2002003542 | A2 | 20030528 | HU 2002-3542 | 20000505 <-- |
| NZ 517237 | A | 20040227 | NZ 2000-517237 | 20000505 |
| AT 305782 | T | 20051015 | AT 2000-931906 | 20000505 |

| | | | | |
|------------------------|----|----------|----------------|----------------------------|
| ES 2249270 | T3 | 20060401 | ES 2000-931906 | 20000505 |
| <u>US 6218418</u> | B1 | 20010417 | US 2000-667603 | 20000922 <-- |
| NO 2002000684 | A | 20020403 | NO 2002-684 | 20020211 <-- |
| HR 2002000128 | A1 | 20030430 | HR 2002-128 | 20020212 <-- |
| MX 2002PA01498 | A | 20030721 | MX 2002-PA1498 | 20020212 <-- |
| ZA 2002001511 | A | 20030311 | ZA 2002-1511 | 20020222 <-- |
| BG 106480 | A | 20020930 | BG 2002-106480 | 20020305 <-- |
| US 7034049 | B1 | 20060425 | US 2002-48486 | 20020501 <-- |
| PRIORITY APPLN. INFO.: | | | | US 1999-372831 A 19990812 |
| | | | | US 2000-560400 A1 20000428 |
| | | | | WO 2000-US6699 W 20000505 |

OTHER SOURCE(S): MARPAT 134:178552
GI

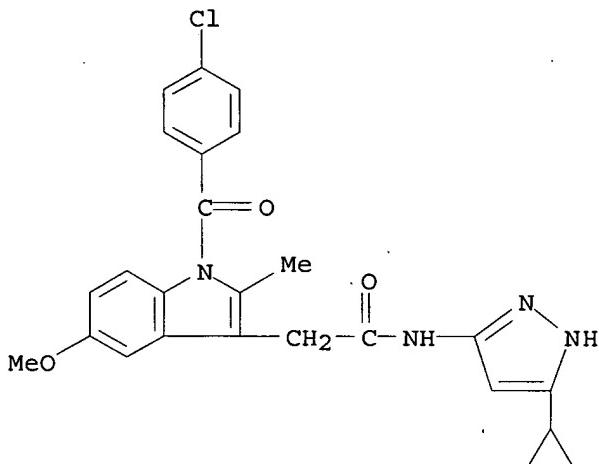


- AB Compds. which are 3-acylaminopyrazole derivs. (I; e.g. N-(5-cyclopropyl-1H-pyrazol-3-yl)-2,2-diphenylacetamide) wherein R is C3-C6 cycloalkyl group optionally substituted by a straight or branched C1-C6 alkyl or arylalkyl group; R1 is a straight or branched C1-C6 alkyl, C2-C4 alkenyl, cycloalkyl, cycloalkenyl, heterocyclyl, aryl, arylalkyl, arylcarbonyl, aryloxalkyl or arylalkenyl group, each of which may be optionally further substituted as indicated in the description; or a pharmaceutically acceptable salt thereof, processes for their preparation and their therapeutic uses. The compds. are useful for the treatment of cancer, cell proliferative disorders, Alzheimer's disease, viral infections, auto-immune diseases or neurodegenerative diseases, but no quant. test results are presented. The cancer is selected from carcinoma, squamous cell carcinoma, hematopoietic tumors of myeloid or lymphoid lineage, tumors of mesenchymal origin, tumors of the central and peripheral nervous system, melanoma, seminoma, teratocarcinoma, osteosarcoma, xeroderma pigmentosum, keratoacanthoma, thyroid follicular cancer and Kaposi's sarcoma. The cell proliferative disorder is selected from benign prostate hyperplasia, familial adenomatosis polyposis, neuro-fibromatosis, psoriasis, vascular smooth cell proliferation associated with atherosclerosis, pulmonary fibrosis, arthritis glomerulonephritis and post-surgical stenosis and restenosis. The method of treatment provides tumor angiogenesis and metastasis inhibition, cell cycle inhibition or cdk/cyclin dependent inhibition, and treatment or prevention of radiotherapy-induced or chemotherapy-induced alopecia. A process for preparing the 3-aminopyrazole derivative or the pharmaceutically acceptable salt thereof, comprising: (a) reacting RCO₂R₂ (R₂ = alkyl), with MeCN in the presence of a basic agent, to obtain RC(O)CH₂CN; (b) reacting RC(O)CH₂CN with hydrazine hydrate to obtain an 3-amino-5-R-1H-pyrazole; (c) oxidizing the 3-amino-5-R-1H-pyrazole to obtain the nitro analog; (d) reacting the nitro compound with tert-butoxycarbonyl anhydride (Boc₂O) to obtain the N-Boc derivative; (e) reducing this BOC derivative to obtain the amino analog; (f) reacting this amino compound with R₁C(O)X (X = OH or a suitable leaving group) to obtain the N1-Boc-protected I; and (g) hydrolyzing this intermediate in an acidic medium to obtain I. Other methods of preparation are also claimed.
- IT 326824-45-7P, 2-[1-(4-Chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]-N-(5-cyclopropyl-1H-pyrazol-3-yl)acetamide
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);

BIOL (Biological study); PREP (Preparation); USES (Uses)
 (acylaminopyrazole derivs., process for preparation and use as antitumor
 agents)

RN 326824-45-7 HCAPLUS

CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-N-(5-cyclopropyl-1H-pyrazol-3-
 yl)-5-methoxy-2-methyl- (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 7 OF 10 HCAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2000:314687 HCAPLUS

DOCUMENT NUMBER: 132:334454

TITLE: Preparation of 2-amino-thiazole derivatives as
 antitumor agents

INVENTOR(S): Pevarello, Paolo; Amici, Raffaella; Traquandi,
 Gabriella; Villa, Manuela; Vulpetti, Anna; Isacchi,
 Antonella

PATENT ASSIGNEE(S): Pharmacia & Upjohn S.p.A., Italy

SOURCE: PCT Int. Appl., 115 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

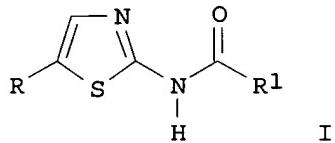
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|--------------|
| WO 2000026202 | A1 | 20000511 | WO 1999-EP8306 | 19991027 <-- |
| W: AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GD, GE, HR, HU, ID,
IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO,
NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2347188 | A1 | 20000511 | CA 1999-2347188 | 19991027 <-- |
| AU 200012679 | A | 20000522 | AU 2000-12679 | 19991027 <-- |
| AU 766193 | B2 | 20031009 | | |
| EP 1124810 | A1 | 20010822 | EP 1999-955931 | 19991027 <-- |

| | | | | |
|--|----|----------|------------------|--------------|
| EP 1124810 | B1 | 20050504 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO | | | | |
| BR 9914958 | A | 20011218 | BR 1999-14958 | 19991027 <-- |
| HU 2001004200 | A2 | 20020328 | HU 2001-4200 | 19991027 <-- |
| HU 2001004200 | A3 | 20031229 | | |
| JP 2002528537 | T | 20020903 | JP 2000-579591 | 19991027 <-- |
| NZ 510965 | A | 20031031 | NZ 1999-510965 | 19991027 <-- |
| TW 222447 | B | 20041021 | TW 1999-88118558 | 19991027 |
| AT 294785 | T | 20050515 | AT 1999-955931 | 19991027 |
| PT 1124810 | T | 20050930 | PT 1999-955931 | 19991027 |
| ES 2241338 | T3 | 20051016 | ES 1999-955931 | 19991027 |
| ZA 2001002870 | A | 20011010 | ZA 2001-2870 | 20010406 <-- |
| NO 2001002057 | A | 20010628 | NO 2001-2057 | 20010426 <-- |
| US 7037929 | B1 | 20060502 | US 2001-807962 | 20010426 <-- |
| MX 2001PA04278 | A | 20020621 | MX 2001-PA4278 | 20010427 <-- |
| IN 2001CN00744 | A | 20050304 | IN 2001-CN744 | 20010528 |
| AU 2004200096 | A1 | 20040205 | AU 2004-200096 | 20040109 |
| PRIORITY APPLN. INFO.: | | | GB 1998-23871 | A 19981030 |
| | | | US 1998-823871 | A 19981030 |
| | | | AU 2000-12679 | A 19991027 |
| | | | WO 1999-EP8306 | W 19991027 |

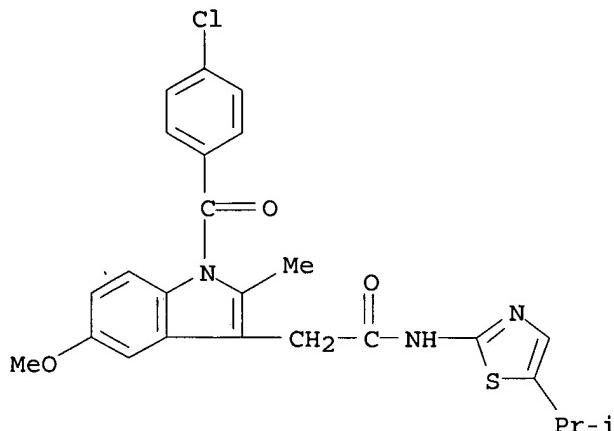
OTHER SOURCE(S) : MARPAT 132:334454
GI



AB The title compds. [I; R = halo, NO₂, (un)substituted amino NH₂, etc.; R₁ = alkyl, alkenyl, 3-6 membered carbocycle, etc.], useful for treating cell proliferative disorders associated with an altered cell dependent kinase activity such as cancer, Alzheimer's disease, viral infections, autoimmune diseases or neurodegenerative disorders, were prepared E.g., thiazole I [R = iso-Pr; R₁ = 4-Me₂NC₆H₄CH₂]. showed Ki of 0.1 μM against cdk2/cyclin A complex.

IT 267656-89-3P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of 2-amino-thiazole derivs. as antitumor agents)

RN 267656-89-3 HCAPLUS
CN 1H-Indole-3-acetamide, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-N-[5-(1-methylethyl)-2-thiazolyl]- (CA INDEX NAME)



REFERENCE COUNT: 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 8 OF 10 HCPLUS COPYRIGHT 2008 ACS on STN
 ACCESSION NUMBER: 1995:858705 HCPLUS
 DOCUMENT NUMBER: 123:266118
 TITLE: Codrugs as a method of controlled drug delivery
 INVENTOR(S): Ashton, Paul; Crooks, Peter Anthony; Riggs, Robert
 Mack; Cynkowski, Tadeusz; Cynkowska, Grazyna
 PATENT ASSIGNEE(S): University of Kentucky Research Foundation, USA
 SOURCE: PCT Int. Appl., 57 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----------------------------------|--|----------|-----------------|--------------|
| WO 9520567 | A1 | 19950803 | WO 1994-US1659 | 19940217 <-- |
| W: AU, CA, JP
RW: AT, BE, CH, | DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE | | | |
| CA 2182228 | A1 | 19950803 | CA 1994-2182228 | 19940217 <-- |
| AU 9462545 | A | 19950815 | AU 1994-62545 | 19940217 <-- |
| AU 705226 | B2 | 19990520 | | |
| EP 740650 | A1 | 19961106 | EP 1994-909643 | 19940217 <-- |
| EP 740650 | B1 | 20040526 | | |
| R: AT, BE, CH, | DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE | | | |
| JP 09509151 | T | 19970916 | JP 1994-520023 | 19940217 <-- |
| AT 267798 | T | 20040615 | AT 1994-909643 | 19940217 |
| PT 740650 | T | 20041029 | PT 1994-909643 | 19940217 |
| ES 2222455 | T3 | 20050201 | ES 1994-909643 | 19940217 |
| US 6051576 | A | 20000418 | US 1997-791071 | 19970129 <-- |
| PRIORITY APPLN. INFO.: | | | US 1994-187462 | A 19940128 |
| | | | WO 1994-US1659 | W 19940217 |
| | | | US 1995-388855 | B1 19950215 |

AB A codrug composition of at least two drug compds. covalently linked to one another via a labile bond to form a single codrug composition, and methods of use of the codrug for the treatment of various medical conditions are disclosed. The codrug may be administered by itself or as a bioerodible or nonbioerodible dosage form, such as injection, liposome, suspension, microsphere, nanoparticle, ointment, transdermal patch, etc.

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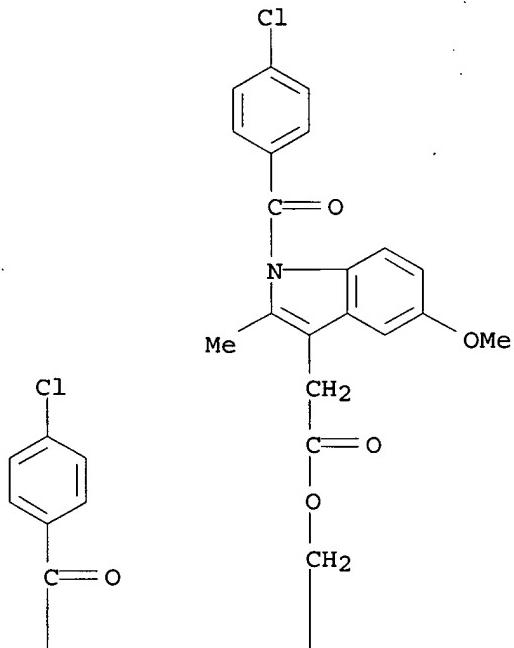
IT 169046-88-2P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(codrug compns. for controlled drug delivery)

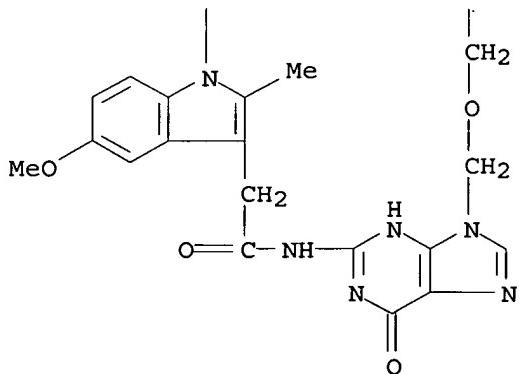
RN 169046-88-2 HCPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-,
2-[[2-[[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indol-3-yl]acetyl]amino]-1,6-dihydro-6-oxo-9H-purin-9-yl]methoxyethyl ester (9CI)
(CA INDEX NAME)

PAGE 1-A



PAGE 2-A

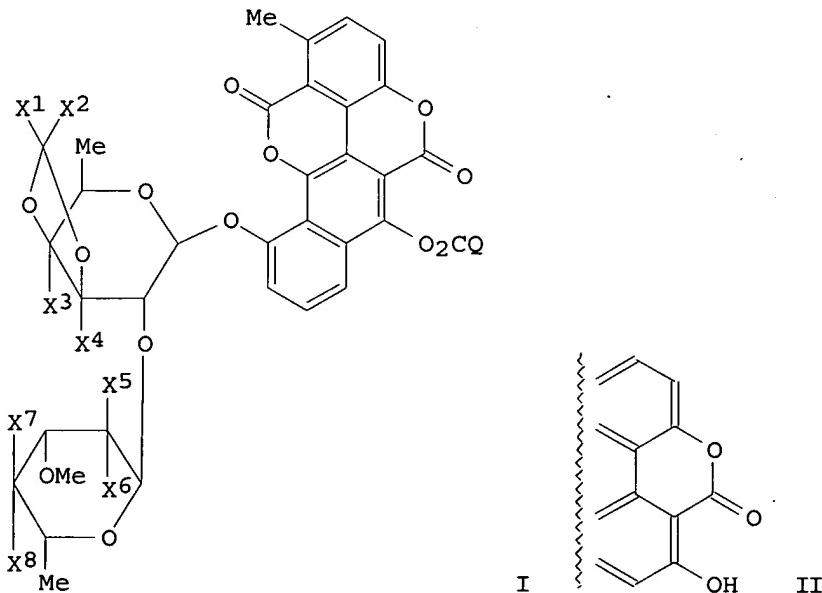


L24 ANSWER 9 OF 10 HCPLUS COPYRIGHT 2008 ACS on STN
ACCESSION NUMBER: 1990:139733 HCPLUS

DOCUMENT NUMBER: 112:139733
 TITLE: Preparation of chartreusin derivatives as antitumor agent and pharmaceutical compositions containing them
 INVENTOR(S): Yamada, Nobutoshi; Sugi, Hideo; Kon, Kenji
 PATENT ASSIGNEE(S): Ishihara Sangyo Kaisha, Ltd., Japan
 SOURCE: Eur. Pat. Appl., 84 pp.
 CODEN: EPXXDW
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| EP 219852 | A2 | 19870429 | EP 1986-114562 | 19861021 <-- |
| EP 219852 | A3 | 19871125 | | |
| EP 219852 | B1 | 19920205 | | |
| R: AT, BE, CH, DE, FR, GB, IT, LI, NL, SE | | | | |
| AT 72447 | T | 19920215 | AT 1986-114562 | 19861021 <-- |
| JP 62174096 | A | 19870730 | JP 1986-251154 | 19861022 <-- |
| CA 1294614 | C | 19920121 | CA 1986-521101 | 19861022 <-- |
| US 4927919 | A | 19900522 | US 1986-922433 | 19861023 <-- |
| US 5064945 | A | 19911112 | US 1989-427370 | 19891027 <-- |
| PRIORITY APPLN. INFO.: | | | | |
| | | | JP 1985-236833 | A 19851023 |
| | | | EP 1986-114562 | A 19861021 |
| | | | US 1986-922433 | A3 19861023 |

OTHER SOURCE(S): CASREACT 112:139733; MARPAT 112:139733
 GI



AB The title compds. [I; X1 = H, (substituted) alkyl; X2 = (substituted) alkyl, (substituted) alkylcarbonylalkyl, etc.; X3, X4 = H, Me; X5 = H, OH, NH2; X6 = H, OH; or X5X6 = O; X7, X8 = H, OH; Q = (substituted) alkyl, (substituted) alkenyl, aminoalkyl, etc.], useful as antitumor agents, are

prepared via reacting alcs. II (X1-X8 same as defined above) with a reactive derivative of HO₂CQ (Q as defined above) in the presence of a condensing agent. Chartreusin was condensed with PhCH(OMe)₂ in CHCl₃ containing p-MeC₆H₄SO₃H and mol. sieve A 1/16 to give II (X₁ = Ph, X₂ = X₃ = X₄ = X₆ = X₇ = H, X₅ = X₈ = OH). This was then condensed with 2-thiophenecarboxylic acid in the presence of DCC to give I (X₁ = Ph, X₂ = X₄ = X₆ = X₇ = H, X₅ = X₈ = OH, O = 2-thienylcarbonyl) (III). III at 60 mg/kg i.p. 3 times a day showed 41% increase in mean survival time over the control in mice implanted with B-16 melanoma.

IT 123938-10-3P

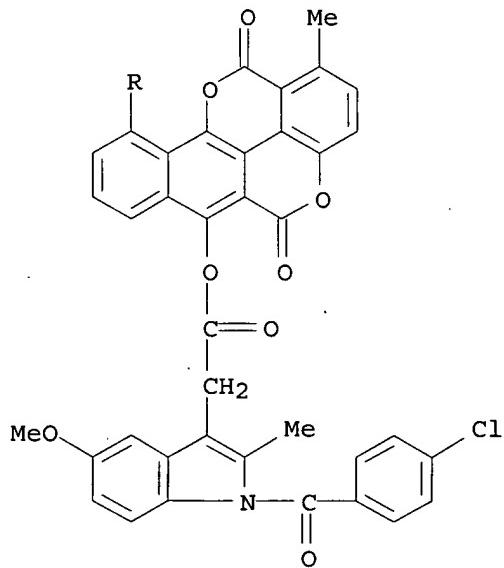
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

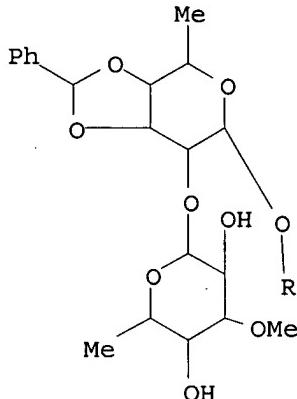
(preparation of, as antitumor agent)

RN 123938-10-3 HCPLUS

CN 1H-Indole-3-acetic acid, 1-(4-chlorobenzoyl)-5-methoxy-2-methyl-, 10-[[6-deoxy-2-O-(6-deoxy-3-O-methyl- α -D-galactopyranosyl)-3,4-O-(phenylmethylene)- β -D-galactopyranosyl]oxy]-5,12-dihydro-1-methyl-5,12-dioxobenzo[h][1]benzopyrano[5,4,3-cde][1]benzopyran-6-yl ester, (R)-(9CI) (CA INDEX NAME)

PAGE 1-A





L24 ANSWER 10 OF 10 HCAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1965:462004 HCAPLUS

DOCUMENT NUMBER: 63:62004

ORIGINAL REFERENCE NO.: 63:11263h,11264a-d

TITLE: Indomethacin antiinflammatory drug

INVENTOR(S): Harman, Robert E.; Kuehl, Frederick A., Jr.; Strachan, Robert G.; Hirschmann, Ralph F.

PATENT ASSIGNEE(S): Merck & Co., Inc.

SOURCE: 12 pp.

DOCUMENT TYPE: Patent

LANGUAGE: Unavailable

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------|-------|----------|-----------------|--------------|
| BE 650447 | ----- | 19650111 | BE 1965-447 | 19640710 <-- |
| US 3285935 | ----- | 19661115 | US 1963-296111 | 19630718 <-- |

PRIORITY APPLN. INFO.:

OTHER SOURCE(S): MARPAT 63:62004

AB Creams and ointments of prolonged action for topical treatment of inflammations are made with the glucuronide (I) of 1-(p-chlorobenzoyl)-2-methyl-5-methoxyindole-3-acetic acid (indomethacin) (II) in an aqueous vehicle or a lower alkyl ester of lower alkanoylated I (III) in an oily or fatty vehicle. Thus, an ointment contains I 1, beeswax 5, anhydrous lanolin 20, heavy mineral oil 5, and white vaseline 31%. Anti-inflammatory action comes from free II slowly liberated from I and III in body tissues by β -glucuronidase therein. Given systemically, I, II, and III are eliminated rapidly by the kidneys (Winter, et al., CA 60, 4660a; Harman, et al., CA 60, 15005c). I is obtained from urine of mammals to which II has been given orally. Thus, a total of 518.5 g. of II-2-14C was given to a rabbit orally with two 1-g. amts. of NH4Cl intraperitoneally at 0 and 12 hrs. and urine of the first 24 hrs. was collected, brought to pH 5, nonconjugated products extracted with C6H6, the extracted urine brought to pH

2.0

and extracted with EtOAc. H2O was added to the exts., EtOAc evaporated and the aqueous residue brought to pH 6.6, and lyophilized to a gum containing 80% of the original radioactivity of the urine. Chromatography on paper with 2:1 MeOH-H2O: 1:1 BuOH-C6H6 revealed the presence of 5-methoxy-2-methylindole-

3-acetic acid (IV) and glucuronides of IV and II. Countercurrent distribution (in a 60 tube train of 30 cc. tubes) of the lyophilized residue in a mixture of 200 cc. aqueous 0.5M Na₂HPO₄ and 200 cc. 0.5M NaH₂PO₄

at

pH 6.6 against 328 cc. EtOAc and 72 cc. sec-BuOH with 75 transfers of 3 cc. each of upper phase per cycle and using scintillation counting plus paper chromatography and uv spectra, located IV near the organic solvent end, the glucuronide of IV at tubes 0-3, and I near tube 29. Tubes 16-34 were combined, aqueous phase separated and brought to pH 2.2 with HCl, recombined

with

the organic phase and 3 extns. made with 15 cc. each of EtOAc. Organic solvent was evaporated and the residue dissolved in H₂O and lyophilized to give 45.5 mg. of I containing 28% of the original radioactivity in the urine. A sample of I is converted 100% into II by glucuronidase. I was converted to the Me ester of the tri-O-acetylglucuronide of III (V) identical with V synthesized from the K salt (VI) of I and tri-O-acetyl- α -D-glucopyranosyl bromide Me uronate (VII). Thus, 5 g. III was suspended in 15 cc. MeOH and 32.8 cc. of 0.427 N K tert-butoxide in tert-BuOH was added dropwise, the solution was dried in vacuo, taken up in acetone, cooled, crysts. filtered, washed, and dried to give 4.2 g. crystals of K salt of II (VIII). VIII (4 g.) was refluxed under N in acetone 2 hrs. with 4.1 g. VII, the solution concentrated to dryness, the residue dissolved in CH₂Cl₂, the solution extracted 3 times with saturated aqueous NaHCO₃, dried over MgSO₄,

and the dry

solution concentrated to a thick yellow oil. The oil was crystallized from ether-n-hexane mixture and gave 2.0 g. of V, m. 150-1°. V was also prepared from crude I from urine by treatment with CH₂N₂ followed by acetylation with Ac₂O.

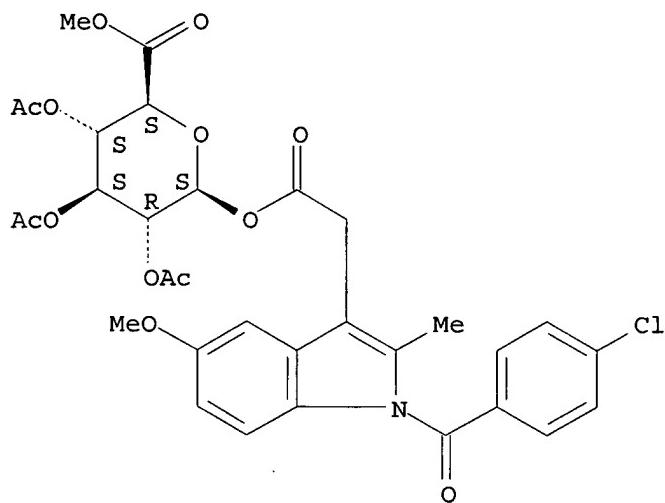
IT 3264-72-0P, Glucuronic acid, methyl ester, triacetate
 1- [1-(p-chlorobenzoyl)-5-methoxy-2-methylindole-3-acetate
 6886-06-2P, Indole-3-acetic acid, 1-(p-chlorobenzoyl)-5-methoxy-2-methyl-, 1-ester with glucuronic acid 7279-30-3P,
 Indole-3-acetic acid, 1-(p-chlorobenzoyl)-5-methoxy-2-methyl-, 1-ester with Me glucuronate triacetate

RL: PREP (Preparation)
 (inflammation-inhibiting preparation containing)

RN 3264-72-0 HCPLUS

CN Glucopyranuronic acid, methyl ester, 2,3,4-triacetate 1-[1-(p-chlorobenzoyl)-5-methoxy-2-methylindole-3-acetate], β -D- (8CI) (CA INDEX NAME)

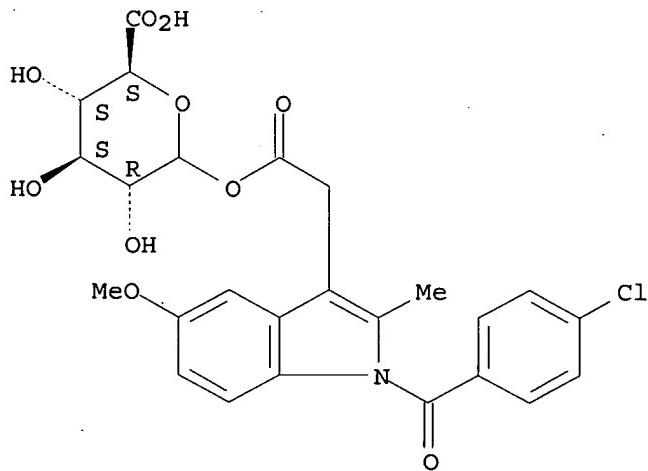
Absolute stereochemistry.



RN 6886-06-2 HCAPLUS

CN D-Glucopyranuronic acid, 1-[1-(4-chlorobenzoyl)-5-methoxy-2-methyl-1H-indole-3-acetate] (CA INDEX NAME)

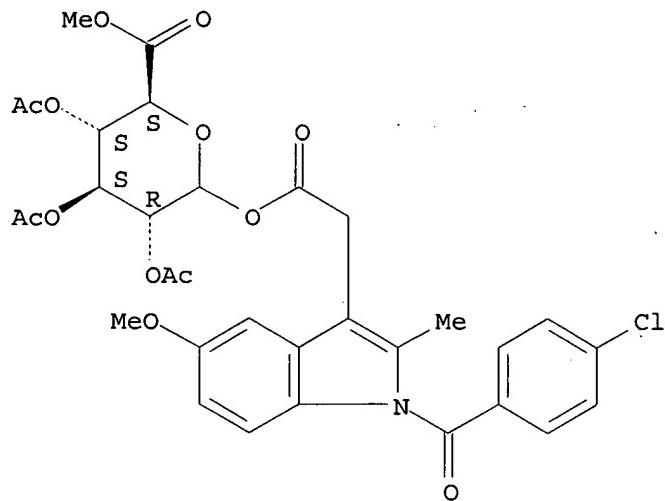
Absolute stereochemistry.



RN 7279-30-3 HCAPLUS

CN Indole-3-acetic acid, 1-(p-chlorobenzoyl)-5-methoxy-2-methyl-, 1-ester with methyl glucuronate triacetate (7CI, 8CI) (CA INDEX NAME)

Relative stereochemistry.



=> log y

COST IN U.S. DOLLARS

SINCE FILE

ENTRY

138.59

TOTAL

SESSION

857.76

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE

ENTRY

-16.00

TOTAL

SESSION

-16.00

CA SUBSCRIBER PRICE

STN INTERNATIONAL LOGOFF AT 07:46:38 ON 07 JAN 2008